Barbara

PTO-1590 (1-2000)

Access DB# <u>700 88</u>

SEARCH REQUEST FORM

Scientific and Technical Information Center $(\mathbb{R}^n_+ + 3)$

Art Unit: 1623 Phon Mail Box and Bldg/Room Locat 8619 (m 1/88) 7 If more than one search is sultable the elected species or structure.	pmitted, please prior ********** he search topic, and descr s, keywords, synonyms, ac ms that may have a specia	Results Format Preferred (circle): PAPER DISK E-MAI ritize searches in order of need. **********************************
Title of Invention:		W
Inventors (please provide full names)	•	
·		
Earliest Priority Filing Date:	10/6/1998	
For Sequence Searches Only Please inc appropriate serial number.	lude all pertinent informatic	on (parent, child, divisional, or issued patent numbers) along with the
Please seam	ah Clamo	Point of Contact: Barb O'Bryen Technical Information Specialist STIC CM1 6A05 308-4291
STAFF USE ONLY	Type of Search	***********
Searcher: ASIB	NA Sequence (#)	Vendors and cost where applicable. STN3//
Searcher Phone #:	AA Sequence (#)	
Searcher Location:	Structure (#)	Questel/Orbit
Date Searcher Picked Up:	Bibliographic	Dr.Link
Date Completed: 7-11-02	Litigation	Lexis/Nexis
Searcher Prep & Review Time: 65	Fulltext	Sequence Systems
Clerical Prep Time:	Patent Family	WWW/Internet
Online Time: 73	Other	Other (specify)

=> fil hcapl FILE 'HCAPLUS' ENTERED AT 10:43:08 ON 11 JUL 2002 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 11 Jul 2002 VOL 137 ISS 2 FILE LAST UPDATED: 10 Jul 2002 (20020710/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

=> d que 131; d que 133; d que 140

```
21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
L1
    (
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L2
L3
             17) SEA FILE=REGISTRY ABB=ON L1 NOT L2
L4
              1) SEA FILE=REGISTRY ABB=ON
                                          57-88-5
L5
             16) SEA FILE=REGISTRY ABB=ON
                                          L3 NOT L4
L6
          14701) SEA FILE=REGISTRY ABB=ON 5 ACETYLOXY
1.7
             15 SEA FILE=REGISTRY ABB=ON L5 NOT L6
          22206 SEA FILE=HCAPLUS ABB=ON FLAVONES+NT,OLD/CT
L11
L12
            458 SEA FILE=HCAPLUS ABB=ON L11(L)?METHOXY?
L13
             41 SEA FILE=HCAPLUS ABB=ON POLYMETHOXYFLAVONE#
L14
           7894 SEA FILE=HCAPLUS ABB=ON
                                          ANTICHOLESTEREMIC AGENTS+OLD/CT
L15
           5890 SEA FILE=HCAPLUS ABB=ON CARDIOVASCULAR AGENTS/CT
L16
           4866 SEA FILE=HCAPLUS ABB=ON
                                          ANTIARTERIOSCLEROTICS/CT
L17
         284065 SEA FILE=HCAPLUS ABB=ON
                                          CARDIOVASCULAR SYSTEM+NT/CT
L18
            777 SEA FILE=HCAPLUS ABB=ON
                                          ?NOBILETIN? OR ?METHYLISOSCUTELLAREIN?
                 OR ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN?
L24
            759 SEA FILE=HCAPLUS ABB=ON
L28
            214 SEA FILE=HCAPLUS ABB=ON
                                          (L24 OR L18 OR L12 OR L13) (L) (BAC OR
                THU OR DMA OR PKT OR PAC)/RL
                                                                               BAC-BIBlogical
L30
          81647 SEA FILE=HCAPLUS ABB=ON
                                         L17(L)(DISEASE# OR DISORDER#)
                                                                                        Activite
L31
             11 SEA FILE=HCAPLUS ABB=ON
                                         L28 AND (L14 OR L15 OR L16 OR L30)
                                                                               THU-therapeutic use
                                                                              DMA - drug machanism
                                                                                        ofaction
L1
             21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
                                                                                PKT- pharmacokinetica
```

1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI PAC-OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/ BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/

```
BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L2
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L3
             17) SEA FILE=REGISTRY ABB=ON
                                          L1 NOT L2
L4
              1) SEA FILE=REGISTRY ABB=ON 57-88-5
L5
             16) SEA FILE=REGISTRY ABB=ON L3 NOT L4
L6
          14701) SEA FILE=REGISTRY ABB=ON
                                         5 ACETYLOXY
L7
             15 SEA FILE=REGISTRY ABB=ON
                                         L5 NOT L6
rac{1}{8}
             21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L9
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L10
              4 SEA FILE=REGISTRY ABB=ON L8 AND L9
L11
          22206 SEA FILE=HCAPLUS ABB=ON FLAVONES+NT, OLD/CT
L12
            458 SEA FILE=HCAPLUS ABB=ON L11(L)?METHOXY?
L13
             41 SEA FILE=HCAPLUS ABB=ON POLYMETHOXYFLAVONE#
L14
           7894 SEA FILE=HCAPLUS ABB=ON ANTICHOLESTEREMIC AGENTS+OLD/CT
L15
           5890 SEA FILE=HCAPLUS ABB=ON
                                        CARDIOVASCULAR AGENTS/CT
L16
           4866 SEA FILE=HCAPLUS ABB=ON
                                        ANTIARTERIOSCLEROTICS/CT
L17
         284065 SEA FILE=HCAPLUS ABB=ON
                                         CARDIOVASCULAR SYSTEM+NT/CT
L18
            777 SEA FILE=HCAPLUS ABB=ON
                                          ?NOBILETIN? OR ?METHYLISOSCUTELLAREIN?
                 OR ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN?
            759 SEA FILE=HCAPLUS ABB=ON
L24
                                         L7
L25
            671 SEA FILE=HCAPLUS ABB=ON
                                         L10
L26
            840 SEA FILE=HCAPLUS ABB=ON
                                         ?TOCOTRIENOL?
L33
              3 SEA FILE=HCAPLUS ABB=ON
                                         (L25 OR L26) AND (L24 OR L18 OR L12
                OR L13) AND (L14 OR L15 OR L16 OR L17)
L1
             21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L2
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L3
             17) SEA FILE=REGISTRY ABB=ON L1 NOT L2
L4
              1) SEA FILE=REGISTRY ABB=ON
                                          57-88-5
L5
             16) SEA FILE=REGISTRY ABB=ON
                                         L3 NOT L4
L6
          14701) SEA FILE=REGISTRY ABB=ON
                                          5 ACETYLOXY
L7
             15 SEA FILE=REGISTRY ABB=ON L5 NOT L6
          22206 SEA FILE=HCAPLUS ABB=ON FLAVONES+NT,OLD/CT
L11
L12
            458 SEA FILE=HCAPLUS ABB=ON
                                         L11(L)?METHOXY?
L13
             41 SEA FILE=HCAPLUS ABB=ON
                                         POLYMETHOXYFLAVONE#
L18
            777 SEA FILE=HCAPLUS ABB=ON
                                         ?NOBILETIN? OR ?METHYLISOSCUTELLAREIN?
                 OR ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN?
L24
            759 SEA FILE=HCAPLUS ABB=ON L7
L34
              1 SEA FILE=REGISTRY ABB=ON CHOLESTEROL/CN
L35
          96409 SEA FILE=HCAPLUS ABB=ON L34 OR CHOLESTEROL/OBI
L36
           2569 SEA FILE=HCAPLUS ABB=ON APOLIPOPROTEIN B/OBI
L37
           8528 SEA FILE=HCAPLUS ABB=ON LOW DENSITY(A)LIPOPROTEIN#/OBI
              8 SEA FILE=HCAPLUS ABB=ON (L24 OR L18 OR L12 OR L13) AND (L35
L38
                OR L36 OR L37)
L40
              6 SEA FILE=HCAPLUS ABB=ON L38 AND PHARMAC?/SC
=> s 131 or 133 or 140
```

L121

13 L31 OR L33 OR L40

Owens 09/528488 Page 3

=> fil wpids

FILE 'WPIDS' ENTERED AT 10:43:11 ON 11 JUL 2002 COPYRIGHT (C) 2002 THOMSON DERWENT

FILE LAST UPDATED: 09 JUL 2002 <20020709/UP>
MOST RECENT DERWENT UPDATE 200243 <200243/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

- >>> Update 2002-42 does not contain any new polymer indexing <<<
- >>> The BATCH option for structure searches has been
 enabled in WPINDEX/WPIDS and WPIX >>>
- >>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY >>>
- >>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://www.derwent.com/dwpi/updates/dwpicov/index.html <<<
- >>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE,
 PLEASE VISIT:
 http://www.stn-international.de/training_center/patents/stn guide.pdf <<</pre>
- >>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER
- GUIDES, PLEASE VISIT:
 http://www.derwent.com/userguides/dwpi_guide.html <<<
- => d que 171; d que 172; s 171 or 172

L42	83	SEA FILE=WPIDS ABB=ON ?NOBILETIN? OR ?METHYLISOSCUTELLAREIN?
L43	3	OR ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN? SEA FILE=WPIDS ABB=ON POLYMETHOXYFLAVONE# OR (POLYMETHOXY OR POLY METHOXY) (W) FLAVONE#
L45	192	SEA FILE=WPIDS ABB=ON ?TOCOTRIENOL?
L46	14714	SEA FILE=WPIDS ABB=ON ?CHOLESTER?
L47	116	SEA FILE=WPIDS ABB=ON (APOLIPOPROTEIN OR APO LIPO PROTEIN OR
		APOLIPO PROTEIN) (W) B
L48	875	SEA FILE=WPIDS ABB=ON LOW DENSITY(W)(LIPOPROTEIN# OR LIPO
		PROTEIN#)
L49	13188	SEA FILE=WPIDS ABB=ON (CARDIOVASCULAR OR CARDIO VASCULAR OR
		HEART OR CARDIAC) (5A) (DISEASE# OR DISORDER#)
L68	12973	SEA FILE=WPIDS ABB=ON ?ARTERIOSCLERO? OR ?ATHEROSCLERO?
L71	3	SEA FILE=WPIDS ABB=ON (L42 OR L43) AND L45 AND ((L46 OR L47
		OR L48 OR L49) OR L68)

L42	83	SEA FILE=WPIDS ABB=ON ?NOBILETIN? OR ?METHYLISOSCUTELLAREIN?
L43		OR ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN? SEA FILE=WPIDS ABB=ON POLYMETHOXYFLAVONE# OR (POLYMETHOXY OR
	_	POLY METHOXY OR POLY METH OXY) (W) FLAVONE#
L45	192	SEA FILE=WPIDS ABB=ON ?TOCOTRIENOL?
L46	14714	SEA FILE=WPIDS ABB=ON ?CHOLESTER?
L47	116	SEA FILE=WPIDS ABB=ON (APOLIPOPROTEIN OR APO LIPO PROTEIN OR
		APOLIPO PROTEIN) (W) B
L48	875	SEA FILE=WPIDS ABB=ON LOW DENSITY(W)(LIPOPROTEIN# OR LIPO
		PROTEIN#)
L49	13188	SEA FILE=WPIDS ABB=ON (CARDIOVASCULAR OR CARDIO VASCULAR OR
	•	HEART OR CARDIAC) (5A) (DISEASE# OR DISORDER#)
L68	12973	SEA FILE=WPIDS ABB=ON ?ARTERIOSCLERO? OR ?ATHEROSCLERO?
L69	11	SEA FILE=WPIDS ABB=ON (L42 OR L43) AND ((L46 OR L47 OR L48 OR
		L49) OR L68)

Owens 09/528488 Page 4

```
L71 3 SEA FILE=WPIDS ABB=ON (L42 OR L43) AND L45 AND ((L46 OR L47 OR L48 OR L49) OR L68)
L72 8 SEA FILE=WPIDS ABB=ON L69 NOT L71
```

L122 11 L71 OR L72

=> fil medl

FILE 'MEDLINE' ENTERED AT 10:43:17 ON 11 JUL 2002

FILE LAST UPDATED: 10 JUL 2002 (20020710/UP). FILE COVERS 1958 TO DATE.

On June 9, 2002, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2002 vocabulary. Enter HELP THESAURUS for details.

THIS FILE CONTAINS CAS REGISTRY NUMBERS FOR EASY AND ACCURATE SUBSTANCE IDENTIFICATION.

=> d que 190; d que 196; s 190 or 196

```
21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
L1 (
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L2
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L3
             17) SEA FILE=REGISTRY ABB=ON L1 NOT L2
L4
              1) SEA FILE=REGISTRY ABB=ON 57-88-5
L5
             16) SEA FILE=REGISTRY ABB=ON L3 NOT L4
L6
          14701) SEA FILE=REGISTRY ABB=ON 5 ACETYLOXY
L7
             15 SEA FILE=REGISTRY ABB=ON
                                         L5 NOT L6
\Gamma8
             21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L9
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L10
              4 SEA FILE=REGISTRY ABB=ON L8 AND L9
L73
           2972 SEA FILE=MEDLINE ABB=ON CARDIOVASCULAR AGENTS/CT
L74
          58947 SEA FILE=MEDLINE ABB=ON ARTERIOSCLEROSIS+NT/CT
L75
           6009 SEA FILE=MEDLINE ABB=ON ANTICHOLESTEREMIC AGENTS/CT
         630810 SEA FILE=MEDLINE ABB=ON A7./CT = candiovascular system
L76
L77
           5731 SEA FILE=MEDLINE ABB=ON APOLIPOPROTEINS B/CT
L78
          15793 SEA FILE=MEDLINE ABB=ON HYPERCHOLESTEROLEMIA+NT/CT
L79
          81692 SEA FILE=MEDLINE ABB=ON CHOLESTEROL+NT/CT
          22993 SEA FILE=MEDLINE ABB=ON LIPOPROTEINS, LDL+NT/CT
T80
           7809 SEA FILE=MEDLINE ABB=ON LIPOPROTEINS, VLDL+NT/CT
L81
L83
            198 SEA FILE=MEDLINE ABB=ON
                                        ?TOCOTRIENOL?
L84
          12234 SEA FILE=MEDLINE ABB=ON
                                         FLAVONES+NT/CT
L85
            114 SEA FILE=MEDLINE ABB=ON
                                         ?NOBILETIN? OR ?METHYLISOSCUTELLAREIN?
                 OR ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN?
L86
              5 SEA FILE=MEDLINE ABB=ON POLYMETHOXYFLAVONE# OR POLYALKYLOXYFLA
                VONE#
L88
             48 SEA FILE=MEDLINE ABB=ON
                                         L7
L89
             41 SEA FILE=MEDLINE ABB=ON
                                         L10
L90
              1 SEA FILE=MEDLINE ABB=ON
                                         (L73 OR L74 OR L75 OR L76 OR L77 OR
                L78 OR L79 OR L80 OR L81) AND (L83 OR L89) AND ((L84 OR L85 OR
```

Page 5

L86) OR L88)

```
L1
             21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L2
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L3
             17) SEA FILE=REGISTRY ABB=ON
                                          L1 NOT L2
L4
              1) SEA FILE=REGISTRY ABB=ON
                                          57-88-5
L5
             16) SEA FILE=REGISTRY ABB=ON
                                          L3 NOT L4
L6
          14701) SEA FILE=REGISTRY ABB=ON
                                          5 ACETYLOXY
L7
             15 SEA FILE=REGISTRY ABB=ON L5 NOT L6
L73
           2972 SEA FILE=MEDLINE ABB=ON CARDIOVASCULAR AGENTS/CT
L74
          58947 SEA FILE=MEDLINE ABB=ON ARTERIOSCLEROSIS+NT/CT
L75
           6009 SEA FILE=MEDLINE ABB=ON ANTICHOLESTEREMIC AGENTS/CT
L76
         630810 SEA FILE=MEDLINE ABB=ON A7./CT
L77
           5731 SEA FILE=MEDLINE ABB=ON APOLIPOPROTEINS B/CT
L78
          15793 SEA FILE=MEDLINE ABB=ON HYPERCHOLESTEROLEMIA+NT/CT
L79
          81692 SEA FILE=MEDLINE ABB=ON CHOLESTEROL+NT/CT
          22993 SEA FILE=MEDLINE ABB=ON LIPOPROTEINS, LDL+NT/CT
L80
          7809 SEA FILE=MEDLINE ABB=ON LIPOPROTEINS, VLDL+NT/CT
L81
L84
                                         FLAVONES+NT/CT
          12234 SEA FILE=MEDLINE ABB=ON
L85
            114 SEA FILE=MEDLINE ABB=ON
                                         ?NOBILETIN? OR ?METHYLISOSCUTELLAREIN?
                 OR ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN?
L86
              5 SEA FILE=MEDLINE ABB=ON POLYMETHOXYFLAVONE# OR POLYALKYLOXYFLA
                VONE#
L88
             48 SEA FILE=MEDLINE ABB=ON L7
L93
            244 SEA FILE=MEDLINE ABB=ON
                                        ?METHOXYFLAVONE?
L96
              5 SEA FILE=MEDLINE ABB=ON (L73 OR L74 OR L75 OR L76 OR L77 OR
                L78 OR L79 OR L80 OR L81) AND ((L85 OR L86) OR L88 OR L93) AND
                L84
```

L123 6 L90 OR L96

=> fil embase

FILE 'EMBASE' ENTERED AT 10:43:22 ON 11 JUL 2002 COPYRIGHT (C) 2002 Elsevier Science B.V. All rights reserved.

FILE COVERS 1974 TO 8 Jul 2002 (20020708/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

 \cdot This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d que 1112; d que 1119

```
L1 ( 21)SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR 1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)

L2 ( 2347)SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL 17)SEA FILE=REGISTRY ABB=ON L1 NOT L2 1)SEA FILE=REGISTRY ABB=ON 57-88-5
```

```
L5
             16) SEA FILE=REGISTRY ABB=ON L3 NOT L4
L6
          14701) SEA FILE=REGISTRY ABB=ON 5 ACETYLOXY
L7
             15 SEA FILE=REGISTRY ABB=ON L5 NOT L6
\Gamma8
             21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L9
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L10
              4 SEA FILE=REGISTRY ABB=ON L8 AND L9
L105
            247 SEA FILE=EMBASE ABB=ON ?TOCOTRIENOL?
            199 SEA FILE=EMBASE ABB=ON L10
L106
L107
            123 SEA FILE=EMBASE ABB=ON L7.
L109
            242 SEA FILE=EMBASE ABB=ON ?NOBILETIN? OR ?CUTELLAREIN? OR
                ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN?
L110
            413 SEA FILE=EMBASE ABB=ON ?METHOXYFLAVONE?
L112
              O SEA FILE=EMBASE ABB=ON (L105 OR L106) AND (L107 OR L109 OR
                L110)
L1 (
             21) SEA FILE=REGISTRY ABB=ON (1178-24-1/BI OR 1244-78-6/BI OR
                1245-15-4/BI OR 1247-97-8/BI OR 14101-61-2/BI OR 1486-56-2/BI
                OR 14965-12-9/BI OR 1721-51-3/BI OR 2174-59-6/BI OR 21763-80-4/
                BI OR 2306-27-6/BI OR 25612-59-3/BI OR 478-01-3/BI OR 481-53-8/
                BI OR 57-88-5/BI OR 57528-78-6/BI OR 6601-66-7/BI OR 6829-55-6/
                BI OR 7678-40-2/BI OR 7741-47-1/BI OR 95943-97-8/BI)
L2 (
           2347) SEA FILE=REGISTRY ABB=ON BENZOPYRAN-6-OL
L3 (
             17) SEA FILE=REGISTRY ABB=ON L1 NOT L2
L4
              1) SEA FILE=REGISTRY ABB=ON 57-88-5
L5
             16) SEA FILE=REGISTRY ABB=ON L3 NOT L4
L6
          14701) SEA FILE=REGISTRY ABB=ON 5 ACETYLOXY
L7
             15 SEA FILE=REGISTRY ABB=ON L5 NOT L6
L97
           9453 SEA FILE=EMBASE ABB=ON CARDIOVASCULAR AGENT/CT OR CARDIAC
                AGENT/CT OR ANTILIPEMIC AGENT/CT OR HYPOCHOLESTEROLEMIC
                AGENT/CT
1,98
          51973 SEA FILE=EMBASE ABB=ON ARTERIOSCLEROSIS+NT/CT
L100
           5789 SEA FILE=EMBASE ABB=ON APOLIPOPROTEIN B/CT
T_1101
          16686 SEA FILE=EMBASE ABB=ON HYPERCHOLESTEROLEMIA+NT/CT
L102
          61370 SEA FILE=EMBASE ABB=ON CHOLESTEROL+NT/CT
L103
          16025 SEA FILE=EMBASE ABB=ON LOW DENSITY LIPOPROTEIN/CT
L104
           6085 SEA FILE=EMBASE ABB=ON VERY LOW DENSITY LIPOPROTEIN/CT
L107
            123 SEA FILE=EMBASE ABB=ON L7
L108
         917224 SEA FILE=EMBASE ABB=ON CARDIOVASCULAR DISEASE+NT/CT
L109
            242 SEA FILE=EMBASE ABB=ON
                                         ?NOBILETIN? OR ?CUTELLAREIN? OR
                ?SINENSETIN? OR ?QUERCITIN? OR ?TANGERETIN?
L110
            413 SEA FILE=EMBASE ABB=ON ?METHOXYFLAVONE?
L117
             26 SEA FILE=EMBASE ABB=ON (L107 OR L109 OR L110) AND (L108 OR
                L97 OR L98 OR (L100 OR L101 OR L102 OR L103 OR L104))
L119
             13 SEA FILE=EMBASE ABB=ON L117 AND (PD OR DT OR PC)/CT
                                                      Subtractings - PD · pharmacology
DT - drug thurspy
PC - prevention
=> dup rem 1123,1121,1119,1122
FILE 'MEDLINE' ENTERED AT 10:43:47 ON 11 JUL 2002
FILE 'HCAPLUS' ENTERED AT 10:43:47 ON 11 JUL 2002
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
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Owens 09/528488 Page 7

FILE 'WPIDS' ENTERED AT 10:43:47 ON 11 JUL 2002

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PROCESSING COMPLETED FOR L123 PROCESSING COMPLETED FOR L121 PROCESSING COMPLETED FOR L119 PROCESSING COMPLETED FOR L122

L124 39 DUP REM L123 L121 L119 L122 (4 DUPLICATES REMOVED)

ANSWERS '1-6' FROM FILE MEDLINE ANSWERS '7-19' FROM FILE HCAPLUS ANSWERS '20-32' FROM FILE EMBASE ANSWERS '33-39' FROM FILE WPIDS

=> d iall 1-6; d ibib abs hitstr 7-19; d iall 20-32; d ibib ab 33-39; fil hom

L124 ANSWER 1 OF 39 MEDLINE

ACCESSION NUMBER: 2001108968 MEDLINE

DOCUMENT NUMBER: 21065691 PubMed ID: 11137857

TITLE: Inhibitory effect of pentalenolactone on vascular smooth

muscle cell proliferation.

AUTHOR: Ikeda M; Fukuda A; Takagi M; Morita M; Shimada Y CORPORATE SOURCE: Department of Veterinary Pharmacology, Faculty of

Agriculture, Miyazaki University, 1-1 Gakuenkibanadai-

nishi, 889-2192, Miyazaki, Japan.

SOURCE: EUROPEAN JOURNAL OF PHARMACOLOGY, (2001 Jan 5) 411 (1-2)

45-53.

Journal code: 1254354. ISSN: 0014-2999.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200102

ENTRY DATE: Entered STN: 20010322

Last Updated on STN: 20010322 Entered Medline: 20010208

ABSTRACT:

The effect of pentalenolactone, an inhibitor of glyceraldehyde-3-phosphate dehydrogenase, on rat vascular smooth muscle cell proliferation was studied. Addition of pentalenolactone together with serum to quiescent cells dose-dependently inhibited cell proliferation and DNA synthesis. This inhibition was not associated with cell death. When quiescent cells were stimulated with serum and then treated with pentalenolactone, the inhibitory effect on the DNA synthesis declined gradually. A similar result was obtained when PD 98059 (2'-amino-3'-methoxyflavone), an inhibitor of extracellular signal-regulated kinase1/2 (ERK1/2) kinase (MEK1/2), was added to the cells after serum stimulation. Pentalenolactone inhibited serum or protein kinase C activator (phorbol 12,13-dibutyrate)-induced phosphorylation of ERK1/2 and MEK1/2. In contrast, pentalenolactone had little effect on platelet-derived growth factor receptor autophosphorylation. Taken together, these results indicate that pentalenolactone inhibits vascular smooth muscle cell proliferation, and that this inhibition appears to be mediated by inhibition of the ERK1/2 cascade.

CONTROLLED TERM: Check Tags: Animal

3T3 Cells

*Antibiotics: PD, pharmacology

Ca(2+)-Calmodulin Dependent Protein Kinase: AI,

antagonists & inhibitors

*Cell Division: DE, drug effects Cell Movement: DE, drug effects

Cells, Cultured

Cyclin-Dependent Kinases: AI, antagonists & inhibitors

DNA: BI, biosynthesis

Owens 09/528488 DNA: DE, drug effects Dose-Response Relationship, Drug Enzyme Inhibitors: PD, pharmacology Flavones: PD, pharmacology Glyceraldehyde-3-Phosphate Dehydrogenases: AI, antagonists & inhibitors Glycolysis: DE, drug effects Mice Mitogen-Activated Protein Kinase Kinases: DE, drug effects Mitogen-Activated Protein Kinase Kinases: ME, metabolism Mitogen-Activated Protein Kinases: DE, drug effects Mitogen-Activated Protein Kinases: ME, metabolism Muscle, Smooth, Vascular: CY, cytology *Muscle, Smooth, Vascular: DE, drug effects Muscle, Smooth, Vascular: ME, metabolism Phosphorylation: DE, drug effects Protein-Serine-Threonine Kinases: DE, drug effects Protein-Serine-Threonine Kinases: ME, metabolism Protein-Tyrosine Kinase: DE, drug effects Protein-Tyrosine Kinase: ME, metabolism Purines: PD, pharmacology Rats Rats, Sprague-Dawley Receptors, Platelet-Derived Growth Factor: DE, drug Receptors, Platelet-Derived Growth Factor: ME, metabolism *Sesquiterpenes: PD, pharmacology Time Factors Tyrosine: DE, drug effects Tyrosine: ME, metabolism p42 MAP Kinase: DE, drug effects p42 MAP Kinase: ME, metabolism 31501-48-1 (arenaemycin E); 55520-40-6 (Tyrosine); 9007-49-2 (DNA) 0 (Antibiotics); 0 (Cyclin-Dependent Kinases); 0 (Enzyme Inhibitors); 0 (Flavones); 0 (PD 98059); 0 (Purines); 0 (Sesquiterpenes); 0 (olomoucine); EC 1.2.1.-

CAS REGISTRY NO.:

CHEMICAL NAME:

(Glyceraldehyde-3-Phosphate Dehydrogenases); EC 2.7.1.-(MEK1 protein); EC 2.7.1.- (MEK2 protein); EC 2.7.1.-(Mitogen-Activated Protein Kinases); EC 2.7.1.-

(Protein-Serine-Threonine Kinases); EC 2.7.1.112 (Protein-Tyrosine Kinase); EC 2.7.10.- (Ca(2+)-Calmodulin

Dependent Protein Kinase); EC 2.7.10.- (Mitogen-Activated Protein Kinase Kinases); EC 2.7.10.- (extracellular

signal-regulated kinase 1); EC 2.7.10.- (p42 MAP Kinase); EC 2.7.11.- (Receptors, Platelet-Derived Growth Factor)

L124 ANSWER 2 OF 39 MEDLINE

ACCESSION NUMBER: 1999243733 MEDLINE

DOCUMENT NUMBER: 99243733 PubMed ID: 10227146

TITLE: Effect of dietary antioxidants on serum lipid contents and

immunoglobulin productivity of lymphocytes in

Sprague-Dawley rats.

AUTHOR: Kaku S; Yunoki S; Mori M; Ohkura K; Nonaka M; Sugano M;

Yamada K

CORPORATE SOURCE: Department of Food Science and Technology, Faculty of

Agriculture, Kyushu University, Fukuoka, Japan.

SOURCE:

BIOSCIENCE, BIOTECHNOLOGY, AND BIOCHEMISTRY, (1999 Mar) 63

(3) 575-6.

Journal code: 9205717. ISSN: 0916-8451.

PUB. COUNTRY:

DOCUMENT TYPE:

Japan Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English Owens 09/528488 Page 9

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199906

ENTRY DATE:

Entered STN: 19990628

Last Updated on STN: 19990628

Entered Medline: 19990615

ABSTRACT:

Sprague-Dawley rats were fed alpha-tocopherol, tocotrienol, or quercetin to examine their dietary effects on serum lipid contents and immunoglobulin productivity. In tocotrienol or quercetin groups, serum triglyceride was lower than in the none group. Moreover, in the alpha-tocopherol group, serum IgA level and IgA productivity of MLN lymphocytes were high, while in the tocotrienol group, IgM productivity of spleen lymphocytes and IgA, IgG, and IgM productivity of MLN lymphocytes were high. Thus, we suggested each antioxidant had different effects in rats.

CONTROLLED TERM:

Check Tags: Animal; Male

*Antioxidants: PD, pharmacology

Cholesterol: BL, blood

*Diet

Immunoglobulin A: BI, biosynthesis Immunoglobulin G: BI, biosynthesis Immunoglobulin M: BI, biosynthesis *Immunoglobulins: BI, biosynthesis

*Lipids: BL, blood

*Lymphocytes: DE, drug effects Lymphocytes: ME, metabolism Quercetin: PD, pharmacology

Rats

Rats, Sprague-Dawley Spleen: CY, cytology Spleen: DE, drug effects Spleen: ME, metabolism Triglycerides: BL, blood

Vitamin E: AA, analogs & derivatives

Vitamin E: PD, pharmacology

CAS REGISTRY NO.:

117-39-5 (Quercetin); 1406-18-4 (Vitamin E); 57-88-5

(Cholesterol)

CHEMICAL NAME:

0 (Antioxidants); 0 (Immunoglobulin A); 0 (Immunoglobulin G); 0 (Immunoglobulin M); 0 (Immunoglobulins); 0 (Lipids);

0 (Triglycerides)

L124 ANSWER 3 OF 39

MEDLINE ACCESSION NUMBER: 2000068727

MEDLINE

DOCUMENT NUMBER: TITLE:

PubMed ID: 10600174 20068727

Inhibitory effect of quercetin metabolites and their related derivatives on copper ion-induced lipid peroxidation in human low-density lipoprotein.

AUTHOR:

Yamamoto N; Moon J H; Tsushida T; Nagao A; Terao J

CORPORATE SOURCE: SOURCE:

Takeda Food Products. Ltd, Itami, Hyogo, 664-0011, Japan. ARCHIVES OF BIOCHEMISTRY AND BIOPHYSICS, (1999 Dec 15) 372

(2) 347-54.

Journal code: 0372430. ISSN: 0003-9861.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200001

ENTRY DATE:

Entered STN: 20000204

Last Updated on STN: 20000204 Entered Medline: 20000124

ABSTRACT:

To determine the antioxidant activity of dietary quercetin (3,3',4',

5,7-pentahydroxyflavone) in the blood circulation, we measured the inhibitory

Owens 09/528488 Page 10

effect of quercetin metabolites and their related derivatives on copper ion-induced lipid peroxidation of human low-density lipoprotein (LDL). Conjugated quercetin metabolites were prepared from the plasma of rat 1 h after oral administration of quercetin aglycone (40 micromol/rat). The rate of cholesteryl ester hydroperoxide (CE-OOH) accumulation and the rate of alpha-tocopherol consumption in mixtures of LDL solution (0.4 mg/ml) with equal volumes of this preparation were slower than the rates in mixtures of LDL with preparations from control rats. The concentrations of CE-OOH after 2 h oxidation in the mixtures of LDL with preparations of conjugated quercetin metabolites were significantly lower than those in the control preparation. It is therefore confirmed that conjugated quercetin metabolites have an inhibitory effect on copper ion-induced lipid peroxidation in human LDL. Quercetin 7-0-beta-glucopyranoside (Q7G) and rhamnetin (3,3',4', 5-tetrahydroxy-7-***methoxyflavone***) exerted strong inhibition and their effect continued even after complete consumption, similarly to quercetin aglycone. The effect of quercetin 3-0-beta-glucopyranoside (Q3G) did not continue after its complete consumption, indicating that the antioxidant mechanism of quercetin conjugates lacking a free hydroxyl group at the 3-position is different from that of the other quercetin conjugates. The result that 4'-0-beta-glucopyranoside (Q4'G) and isorhamnetin (3,4',5, 7-tetrahydroxy-3'-methoxyflavone) showed little inhibition implies that introduction of a conjugate group to the position of the dihydroxyl group in the B ring markedly decreases the inhibitory effect. The results of azo radical-induced lipid peroxidation of LDL and the measurement of free radical scavenging capacity using stable free radical, 1,1,-diphenyl-2-picrylhydrazyl, demonstrated that the o-dihydroxyl structure in the B ring is required to exert maximum free radical scavenging activity. It is therefore likely that conjugation occurs at least partly in positions other than the B ring during the process of metabolic conversion so that the inhibitory effect of dietary quercetin is retained in blood plasma after absorption.

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CONTROLLED TERM:
                    Check Tags: Animal; Human; Male; Support, Non-U.S. Gov't
                     Amidines: AI, antagonists & inhibitors
                     Amidines: PD, pharmacology
                     Antioxidants: CH, chemistry
                     Antioxidants: ME, metabolism
                     Antioxidants: PD, pharmacology
                     Bepridil: AA, analogs & derivatives
                     Bepridil: ME, metabolism
                       Cholesterol Esters: ME, metabolism
                     Copper Sulfate: AI, antagonists & inhibitors
                    *Copper Sulfate: PD, pharmacology
                     Cysteine: ME, metabolism
                     Free Radical Scavengers: CH, chemistry
                     Free Radical Scavengers: ME, metabolism
                     Free Radical Scavengers: PD, pharmacology
                     Free Radicals: ME, metabolism
                     Kinetics
                    *Lipid Peroxidation: DE, drug effects
                      *Lipoproteins, LDL: ME, metabolism
                     Models, Chemical
                     Oxidants: AI, antagonists & inhibitors
                     Oxidants: PD, pharmacology
                     Oxidation-Reduction: DE, drug effects
                       Quercetin: AA, analogs & derivatives
                       Quercetin: CH, chemistry
                      *Quercetin: ME, metabolism
                      *Quercetin: PD, pharmacology
                     Rats
                     Rats, Wistar
                     Vitamin E: ME, metabolism
CAS REGISTRY NO.:
                    117-39-5 (Quercetin); 13217-66-8 (2,2'-azobis(2-
```

Owens 09/528488 Page 11

amidinopropane)); 1406-18-4 (Vitamin E); 1898-66-4 (2,2-diphenyl-1-picrylhydrazyl); 2058-59-5 (cholesteryl ester hydroperoxide); 52-90-4 (Cysteine); 64706-54-3

(Bepridil); 7758-98-7 (Copper Sulfate)

CHEMICAL NAME: 0 (Amidines); 0 (Antioxidants); 0 (Cholesterol Esters); 0

(Free Radical Scavengers); 0 (Free Radicals); 0

(Lipoproteins, LDL); 0 (Oxidants)

L124 ANSWER 4 OF 39 MEDLINE

ACCESSION NUMBER: 95218768 MEDLINE

DOCUMENT NUMBER: 95218768 PubMed ID: 7703977

TITLE: Cardiotonic flavonoids from Citrus plants (Rutaceae).

AUTHOR: Itoigawa M; Takeya K; Furukawa H

Tokaigakuen Women's College, Nagoya, Japan. CORPORATE SOURCE:

SOURCE: BIOLOGICAL AND PHARMACEUTICAL BULLETIN, (1994 Nov) 17 (11)

1519-21.

Journal code: 9311984. ISSN: 0918-6158.

PUB. COUNTRY: Japan

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199505

Entered STN: 19950518 ENTRY DATE:

> Last Updated on STN: 19980206 Entered Medline: 19950508

ABSTRACT:

Two flavonoids, 3,5,6,7,8,3',4'-heptamethoxyflavone (HEPTA) and natsudaidain isolated from Citrus plants (Rutaceae), produced a positive inotropic effect (PIE) on guinea-pig papillary muscle. Natsudaidain (pD2 4.98 +/- 0.07) was more potent than HEPTA (pD2 4.33 +/- 0.08), but the maximum PIE of HEPTA was greater than that of natsudaidain. The PIE of HEPTA was completely inhibited by reserpinization of the guinea pig, and partially inhibited by metoprolol and carbachol. The carbachol inhibition was omitted by atropine. The mechanism of PIE of HEPTA is accounted for catecholamine release from cardiac tissue.

CONTROLLED TERM: Check Tags: Animal; Female; Male

> Cardiotonic Agents: AD, administration & dosage Cardiotonic Agents: IP, isolation & purification

*Cardiotonic Agents: PD, pharmacology

*Citrus: CH, chemistry

Dose-Response Relationship, Drug

Flavones: AD, administration & dosage Flavones: IP, isolation & purification

*Flavones: PD, pharmacology

Guinea Pigs Methylation

*Myocardial Contraction: DE, drug effects Papillary Muscles: DE, drug effects

Plant Extracts: AD, administration & dosage Plant Extracts: IP, isolation & purification

Plant Extracts: PD, pharmacology Plant Leaves: CH, chemistry

Rats

Structure-Activity Relationship

Time Factors

CAS REGISTRY NO.: 1178-24-1 (3,3',4',5,6,7,8-heptamethoxyflavone);

35154-55-3 (natsudaidain)

CHEMICAL NAME: 0 (Cardiotonic Agents); 0 (Flavones); 0 (Plant Extracts)

L124 ANSWER 5 OF 39 MEDLINE

95131371 ACCESSION NUMBER: MEDLINE

DOCUMENT NUMBER: 95131371 PubMed ID: 7830234 Owens 09/528488 Page 12

TITLE: Anti-invasive activity of 3,7-dimethoxyflavone in

vitro.

AUTHOR: Parmar V S; Jain R; Sharma S K; Vardhan A; Jha A; Taneja P;

Singh S; Vyncke B M; Bracke M E; Mareel M M

CORPORATE SOURCE: I

Department of Chemistry, University of Delhi, India. JOURNAL OF PHARMACEUTICAL SCIENCES, (1994 Sep) 83 (9)

1217-21.

Journal code: 2985195R. ISSN: 0022-3549.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

SOURCE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199502

ENTRY DATE:

Entered STN: 19950307

Last Updated on STN: 19970203 Entered Medline: 19950217

ABSTRACT:

Invasion of MCF-7/6 human mammary carcinoma cells into embryonic chick heart fragments was studied in organ culture during 8 days. The effect of 31 polyphenolic compounds, belonging to the flavonoids, chalcones, or coumarins, was tested in this assay for invasion. The anti-invasive activity of 3,7-***dimethoxyflavone*** was found at concentrations ranging from 1 to 100 microM. At these anti-invasive concentrations, no cytotoxic effects could be detected: the anti-invasive effect was reversible upon omission of the molecule from the medium, and treatment of MCF-7/6 cells or heart fragments did not affect subsequent outgrowth from explants on tissue culture plastic. The molecule did not inhibit growth of MCF-7/6 cell aggregates nor of heart fragments kept in suspension culture. The action mechanism of 3,7-***dimethoxyflavone*** is the subject of our ongoing research.

CONTROLLED TERM:

Check Tags: Animal; Human; Support, Non-U.S. Gov't

*Antineoplastic Agents: PD, pharmacology

Breast Neoplasms Chick Embryo

*Flavones: PD, pharmacology Heart: EM, embryology

*Neoplasm Invasiveness: PA, pathology

Organ Culture

Structure-Activity Relationship

Tumor Cells, Cultured

CHEMICAL NAME:

0 (3,7-dimethoxyflavone); 0 (Antineoplastic

Agents); 0 (Flavones)

L124 ANSWER 6 OF 39 MEDLINE

ACCESSION NUMBER:

89168929 MEDLINE

DOCUMENT NUMBER:

89168929 PubMed ID: 2924447

TITLE:

The flavonoid tangeretin inhibits invasion of MO4 mouse cells into embryonic chick heart in vitro.

AUTHOR:

Bracke M E; Vyncke B M; Van Larebeke N A; Bruyneel E A; De

Bruyne G K; De Pestel G H; De Coster W J; Espeel M F;

Mareel M M

CORPORATE SOURCE:

Department of Radiotherapy and Nuclear Medicine, University

Hospital, Gent, Belgium.

SOURCE:

CLINICAL AND EXPERIMENTAL METASTASIS, (1989 May-Jun) 7 (3)

283-300.

Journal code: 8409970. ISSN: 0262-0898.

PUB. COUNTRY:

ENGLAND: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

198905

ENTRY DATE:

Entered STN: 19900306

Last Updated on STN: 19900306

Owens 09/528488 Page 13

Entered Medline: 19890505

ABSTRACT:

Tangeretin, a flavonoid from citrus plants, was found to inhibit the invasion of MO4 cells (Kirsten murine sarcoma virus transformed fetal mouse cells) into embryonic chick heart fragments in vitro. The flavonoid appeared to be chemically stable in tissue culture medium, and the anti-invasive effect was reversible on omission of the molecule from the medium. Unlike (+)-catechin, another anti-invasive flavonoid, tangeretin bound poorly to extracellular matrix. It did not alter fucosylated surface glycopeptides of MO4 cells. Tangeretin seemed not to act as a microtubule inhibitor, as immunocytochemistry revealed no disturbance of the cytoplasmic microtubule complex. However, at anti-invasive concentrations of tangeretin, cell proliferation and thymidine incorporation appeared to be inhibited. When cultured on an artificial substrate, treated MO4 cells were less elongated, covered a larger surface area and exhibited a slower directional migration than untreated cells. From the decrease in ATP content in MO4 cells after ***tangeretin*** treatment, we deduce that this flavonoid inhibits a number of intracellular processes, which leads to an inhibition of cell motility and hence of invasion.

CONTROLLED TERM: Check Tags: Animal; Support, Non-U.S. Gov't

Adenosine Triphosphate: ME, metabolism

Cell Aggregation

Cell Line

Cell Movement: DE, drug effects

Chick Embryo DNA Replication

*Flavones: PD, pharmacology

Fucose: AN, analysis

Glycopeptides: IP, isolation & purification

*Heart: DE, drug effects

Mice

Mice, Inbred C3H

Microtubules: DE, drug effects Microtubules: UL, ultrastructure

*Myocardium: PA, pathology

*Neoplasm Invasiveness: UL, ultrastructure

Organ Culture

*Sarcoma, Experimental: PA, pathology

Sarcoma, Experimental: PP, physiopathology Sarcoma, Experimental: UL, ultrastructure

CAS REGISTRY NO.: 3713-31-3 (Fucose); **481-53-8 (tangeretin)**;

56-65-5 (Adenosine Triphosphate)

CHEMICAL NAME: 0 (Flavones); 0 (Glycopeptides)

L124 ANSWER 7 OF 39 HCAPLUS COPYRIGHT 2002 ACS DUPLICATE 1

ACCESSION NUMBER: 2001:713074 HCAPLUS

DOCUMENT NUMBER: 135:251964

TITLE: Compositions and methods using

polymethoxyflavones for treating, reducing,

and preventing cardiovascular diseases and disorders

INVENTOR(S): Horowitz, Robert M.; Guthrie, Najla; Kurowska,

Elzbieta Maria; Manthey, John A.

PATENT ASSIGNEE(S): KGK Synergie, Can.; United States Department of

Agriculture

SOURCE: PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                            DATE
                                                            20010316
    WO 2001070029
                       Α1
                            20010927
                                           WO 2001-US8395
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                        US 2000-528488
                                                         A 20000317
    Compns. and methods for the treatment, redn., and/or prevention of
    cardiovascular diseases and disorders are described. Individuals at high
    risk for developing or having cardiovascular disease or disorder may be
    treated with an ED of a polymethoxyflavone including limocitrin
    derivs., quercetin derivs., naturally occurring
    polymethoxyflavones, tocotrienols, and mixts. of these
     compds.
IT
     478-01-3 481-53-8 1178-24-1 1244-78-6
     1245-15-4 1247-97-8 1486-56-2
     1721-51-3, .alpha.-Tocotrienol 2174-59-6
     2306-27-6 6601-66-7 6829-55-6,
     Tocotrienol 7678-40-2 7741-47-1
     14101-61-2, .gamma.-Tocotrienol 14965-12-9
     21763-80-4 25612-59-3, .delta.-Tocotrienol
     95943-97-8
     RL: BAC (Biological activity or effector, except adverse); BSU
     (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (polymethoxyflavones for cardiovascular disease treatment)
RN
     478-01-3 HCAPLUS
     4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7,8-tetramethoxy- (9CI)
CN
       (CA INDEX NAME)
```

RN 481-53-8 HCAPLUS CN 4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 1178-24-1 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3,5,6,7,8-pentamethoxy-(9CI) (CA INDEX NAME)

RN 1244-78-6 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3-hydroxy-5,7-dimethoxy-(9CI) (CA INDEX NAME)

RN 1245-15-4 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-3,7-dimethoxy-(9CI) (CA INDEX NAME)

RN 1247-97-8 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3,5,7-trimethoxy- (9CI) (CA INDEX NAME)

RN 1486-56-2 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-[3,4-bis(phenylmethoxy)phenyl]-3,5-dimethoxy-7-(phenylmethoxy)- (9CI) (CA INDEX NAME)

RN 1721-51-3 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,5,7,8-tetramethyl-2-(4,8,12-trimethyl-3,7,11-tridecatrienyl)- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

= CMe₂

RN 2174-59-6 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-6,7,8-trimethoxy-(9CI) (CA INDEX NAME)

RN 2306-27-6 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7-trimethoxy- (9CI) (CA INDEX NAME)

RN 6601-66-7 HCAPLUS

CN 4H-1-Benzopyran-4-one, 5,7,8-trimethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 6829-55-6 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2-methyl-2-(4,8,12-trimethyl-3,7,11-tridecatrienyl)- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

= CMe₂

RN 7678-40-2 HCAPLUS

CN 4H-1-Benzopyran-4-one, 3-(acetyloxy)-2-(3,4-dimethoxyphenyl)-5,7-dimethoxy-(9CI) (CA INDEX NAME)

RN 7741-47-1 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3,5,7,8-tetramethoxy- (9CI) (CA INDEX NAME)

MeO OMe OMe OMe

RN 14101-61-2 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,7,8-trimethyl-2-[(3E,7E)-4,8,12-trimethyl-3,7,11-tridecatrienyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

RN 14965-12-9 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-3,7,8-trimethoxy-(9CI) (CA INDEX NAME)

RN 21763-80-4 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-6,7-dimethoxy-(9CI) (CA INDEX NAME)

RN 25612-59-3 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,8-dimethyl-2-[(3E,7E)-4,8,12-trimethyl-3,7,11-tridecatrienyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

RN 95943-97-8 HCAPLUS

НО

CN 4H-1-Benzopyran-4-one, 3,5,7-triethoxy-2-(4-ethoxy-3-methoxyphenyl)-8-

methoxy- (9CI) (CA INDEX NAME)

57-88-5, Cholesterol, biological studies ΙT

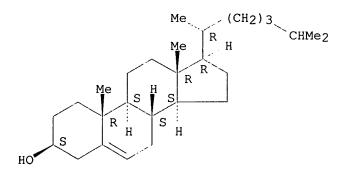
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(polymethoxyflavones for cardiovascular disease treatment)

57-88-5 HCAPLUS RN

CN Cholest-5-en-3-ol (3.beta.)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS 4 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L124 ANSWER 8 OF 39 HCAPLUS COPYRIGHT 2002 ACS DUPLICATE 2

ACCESSION NUMBER:

2001:338337 HCAPLUS

DOCUMENT NUMBER:

134:357559

TITLE:

Modification of cholesterol concentrations

with citrus phytochemicals

INVENTOR(S):

McGill, Carla R.; Green, Nancy R. Tropicana Products, Inc., USA

PATENT ASSIGNEE(S):

SOURCE:

PCT Int. Appl., 34 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

1

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

RU, TJ, TM

PATENT INFORMATION:

PATENT	NO.		KI	ND	DATE			A	PPLI	CATI	ON NO	ο.	DATE			
							_									
WO 2001	0321	60	A	2	2001	0510		W	0 20	00-U	S417	84	2000	1101		
WO 2001032160		A	3	20020321												
₩:	ΑE,	AG,	AL,	AM,	ΑT,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,
	CN,	CR,	CU,	CZ,	CZ,	DE,	DE,	DK,	DK,	DM,	DZ,	EE,	EE,	ES,	FI,	FI,
	GB,	GD,	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,
	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,

Searched by Barb O'Bryen, STIC 308-4291

NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD,

Page 20

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 2002006953 Α1 20020117 US 1999-435304 19991105 PRIORITY APPLN. INFO.: US 1999-435304 A 19991105 Methods, products and compns. are provided which, when administered to a mammal, including humans, raises HDL serum cholesterol levels, while typically also lowering the ratio of LDL to HDL serum cholesterol levels. An effective amt. of one or more of a monoterpene, a terpene and a flavonoid are included in the treatment compn. ΙT 57-88-5D, Cholesterol, HDL conjugates RL: BOC (Biological occurrence); BSU (Biological study, unclassified); MFM (Metabolic formation); THU (Therapeutic use); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence); USES (Uses) (modification of **cholesterol** concns. with citrus phytochems.) 57-88-5 HCAPLUS RN CN Cholest-5-en-3-ol (3.beta.)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Me
$$(CH_2)_3$$
 $CHMe_2$

Me R H S H S H

IT 478-01-3, Nobiletin 481-53-8, Tangeretin 2306-27-6, Sinensetin 7741-47-1

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PROC (Process); USES (Uses) (modification of cholesterol concns. with citrus phytochems.)
478-01-3 HCAPLUS
4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7,8-tetramethoxy- (9CI)

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7,8-tetramethoxy- (9CI (CA INDEX NAME)

RN 481-53-8 HCAPLUS

RN

CN 4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 2306-27-6 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7-trimethoxy- (9CI) (CA INDEX NAME)

RN 7741-47-1 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3,5,7,8-tetramethoxy- (9CI) (CA INDEX NAME)

L124 ANSWER 9 OF 39 HCAPLUS COPYRIGHT 2002 ACS DUPLICATE 3

ACCESSION NUMBER:

2000:240940 HCAPLUS

DOCUMENT NUMBER:

132:260708

TITLE:

Compositions and methods of inhibiting neoplastic and cardiovascular diseases with compounds related to

limocitrin and 5-desmethyl sinensetin

INVENTOR(S):

Guthrie, Najla; Manthey, John A.; Horowitz, Robert M.

PATENT ASSIGNEE(S):

KGK Synergize, Can.; Usda-Ars-Ott

SOURCE:

PCT Int. Appl., 29 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.			KI	ND	DATE			A	PPĻI	CATI	ON NO	o. :	DATE				
WO 2000019998			A	A1 20000413				WO 1999-US23238 19991005									
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	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	
	KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MD,	MG,	MK,	MN,	MW,	
	MX,	NO,	ΝZ,	ΡL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	
	TT,	UA,	UG,	UZ,	VN,	YU,	ZA,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM

RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG AU 9962916 20000426 A1 AU 1999-62916 19991005 EP 1119353 20010801 A1 EP 1999-950209 19991005 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO

PRIORITY APPLN. INFO.: US 1998-167634 A 19981006 WO 1999-US23238 W 19991005

AB Compns. and methods for the prevention and treatment of neoplastic diseases and cardiovascular diseases (e.g. atherosclerosis) are described. Individuals at a high risk of developing or having neoplasia or atherosclerosis undergoing conventional therapies may be treated with an ED of limocitrin compds. including, but not limited to e.g. 3,5,7,4'-tetramethoxylimocitrin, limocitrin and 5-desmethylsinensetin.

TT 7741-47-1P 14965-12-9P 95943-97-8P
RL: BAC (Biological activity or effector, except adverse); BSU
(Biological study, unclassified); SPN (Synthetic preparation); THU
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(limocitrin derivs. and desmethyl **sinensetin** for inhibition of neoplastic and cardiovascular diseases)

RN 7741-47-1 HCAPLUS

CN

4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3,5,7,8-tetramethoxy- (9CI) (CA INDEX NAME)

RN 14965-12-9 HCAPLUS CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-3,7,8-trimethoxy-(9CI) (CA INDEX NAME)

RN 95943-97-8 HCAPLUS

CN 4H-1-Benzopyran-4-one, 3,5,7-triethoxy-2-(4-ethoxy-3-methoxyphenyl)-8methoxy- (9CI) (CA INDEX NAME)

IT 478-01-3, Nobiletin 481-53-8, Tangeretin 1244-78-6 1247-97-8, Quercetin pentamethyl ether 1486-56-2 2174-59-6 2306-27-6, Sinensetin 6601-66-7 21763-80-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (limocitrin derivs. and desmethyl sinensetin for inhibition

(limocitrin derivs. and desmethyl sinensetin for inhibit of neoplastic and cardiovascular diseases)

RN 478-01-3 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7,8-tetramethoxy- (9CI) (CA INDEX NAME)

RN 481-53-8 HCAPLUS CN 4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 1244-78-6 HCAPLUS
CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3-hydroxy-5,7-dimethoxy-(9CI) (CA INDEX NAME)

RN 1247-97-8 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-3,5,7-trimethoxy- (9CI) (CA INDEX NAME)

MeO O OMe OMe

RN 1486-56-2 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-[3,4-bis(phenylmethoxy)phenyl]-3,5-dimethoxy-7-(phenylmethoxy)- (9CI) (CA INDEX NAME)

Ph-CH₂-O.

Ome
Ome
OCH₂-Ph

RN 2174-59-6 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-6,7,8-trimethoxy-(9CI) (CA INDEX NAME)

MeO OMe OMe

RN 2306-27-6 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7-trimethoxy- (9CI) (CA INDEX NAME)

MeO O OMe

RN 6601-66-7 HCAPLUS

CN 4H-1-Benzopyran-4-one, 5,7,8-trimethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 21763-80-4 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-6,7-dimethoxy-(CA INDEX NAME)

REFERENCE COUNT:

5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L124 ANSWER 10 OF 39 HCAPLUS COPYRIGHT 2002 ACS

DUPLICATE 4

ACCESSION NUMBER:

1999:231499 HCAPLUS

DOCUMENT NUMBER:

130:262145

TITLE:

Use of citrus limonoids and flavonoids as well as

tocotrienols for the treatment of cancer and

hypercholesterolemia

INVENTOR(S):

PATENT ASSIGNEE(S):

Carrol, Kenneth Kitchener; Kurowska, Elzbieta Maria KGK Synergize Inc., Can.; Carroll, Margaret Aileen;

Guthrie, Najla

SOURCE:

PCT Int. Appl., 31 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

2

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

P.F	ATENT	NO.		KI	ND	DATE			A	PPLI	CATI	ои ис	ο.	DATE			
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														JP,			
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CF	2304	202		A	Ą	1999	0401		C	A 19	98-2	30420	02	1998	0924		
JΑ	J 9894	557		A.	1	1999	0412		A	U 19	98-9	4557		1998	0924		
ĒΕ	1049	464		A:	2	2000	1108		Ε	P 19	98-9	47740	0	1998	0924		
	R:	ΑT,	DE,	FR,	GB,	ΙT,	NL										
PRIORIT	Y APP	LN.	INFO	.:					US 1	997-	9386	40	Α	1997	0926		
									WO 1	998-	IB17	21	W	1998	0924		

Compns. and methods for the prevention and treatment of neoplastic AB

diseases and hypercholesterolemia are described. Individuals at a high risk of developing or having neoplasia or hypercholesterolemia undergoing conventional therapies may be treated with an ED of triterpene derivs. in citrus limonoids, polyphenolic flavonoid citrus compds.,

tocotrienols or a combination of these agents.

IT 478-01-3, Nobiletin 481-53-8,

Tangeretin 1721-51-3, .alpha.-Tocotrienol

6829-55-6, Tocotrienol 14101-61-2, .gamma.-

Tocotrienol 25612-59-3, .delta.-Tocotrienol

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(citrus limonoids and flavonoids as well as tocotrienols for

treatment of cancer and hypercholesterolemia)

RN 478-01-3 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7,8-tetramethoxy- (9CI)

(CA INDEX NAME)

RN 481-53-8 HCAPLUS

CN 4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 1721-51-3 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,5,7,8-tetramethyl-2-(4,8,12-trimethyl-3,7,11-tridecatrienyl)- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

= CMe₂

RN 6829-55-6 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2-methyl-2-(4,8,12-trimethyl-3,7,11-tridecatrienyl)- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

= CMe $_2$

RN 14101-61-2 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,7,8-trimethyl-2-[(3E,7E)-4,8,12-trimethyl-3,7,11-tridecatrienyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

RN 25612-59-3 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,8-dimethyl-2-[(3E,7E)-4,8,12-trimethyl-3,7,11-tridecatrienyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

IT 57-88-5, Cholesterol, biological studies

57-88-5D, Cholesterol, esters

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

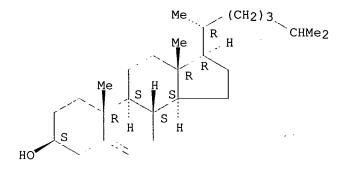
(citrus limonoids and flavonoids as well as tocotrienols for

treatment of cancer and hypercholesterolemia)

· RN 57-88-5 HCAPLUS

CN Cholest-5-en-3-ol (3.beta.) - (9CI) (CA INDEX NAME)

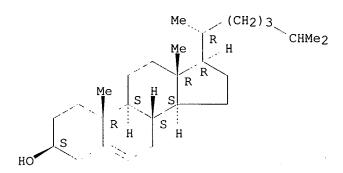
Absolute stereochemistry.



RN 57-88-5 HCAPLUS

CN Cholest-5-en-3-ol (3.beta.)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L124 ANSWER 11 OF 39 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

2001:392055 HCAPLUS

DOCUMENT NUMBER:

135:10008

TITLE:

Compositions and methods for treatment of neoplastic

diseases with combinations of limonoids, flavonoids

and tocotrienols

INVENTOR(S):

Guthrie, Najla; Kurowska, Elzbieta Maria

PATENT ASSIGNEE(S):

KGK Synergize, Can.

SOURCE:

U.S., 7 pp., Cont.-in-part of U.S. Ser. No. 938,640,

abandoned.
CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6239114	В1	20010529	US 2000-481963	20000112
US 6251400	B1	20010626	US 1997-938640	19970926

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WO 2001051043
                          A2
                                20010719
                                                WO 2001-IB186
                                                                    20010112
     WO 2001051043
                          Α3
                                20020530
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              SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
          RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                             US 1997-938640
                                                                B2 19970926
                                             US 2000-481963
                                                                A 20000112
     Compns. and methods for the prevention and treatment of neoplastic
     diseases using a synergistic combination of triterpenes are described.
     Individuals at a high risk of developing or having neoplasia undergoing
     conventional therapies may be treated with an ED of triterpene derivs.,
     i.e., limonoids (1-500 mg/day), flavonoids (200-5000 mg/day),
     tocotrienols (1-1200 mg/day) or a combination of these agents.
     For example, in the DU 145 prostatic tumor cell line, tangeretin
     alone or nobitelin alone inhibited these cells most effectively followed
     by nomilin when the test agents were given alone. When given as
     combinations, the most effective combination was nomilin + hesperitin +
     .alpha.-tocotrienol, followed by limolin + nobelitin + .alpha.-
     tocotrienol and nomilin + naringenin, followed by nomilin +
     hesperitin + .alpha.-tocotrienol and limonin +
     tangeretin + .alpha.-tocopherol, followed by nomilin +
     tangeretin and limonin + tangeretin, followed by limonin
     + naringenin.
TT
     478-01-3, Nobiletin 481-53-8,
     Tangeretin 1721-51-3, .alpha.-Tocotrienol
     6829-55-6, Tocotrienol 14101-61-2, .gamma.-
     Tocotrienol 25612-59-3, .delta.-Tocotrienol
     RL: BAC (Biological activity or effector, except adverse); BSU
     (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
         (compns. of synergistic combinations of limonoids, flavonoids and
         tocotrienols for treatment of neoplastic diseases)
RN
     478-01-3 HCAPLUS
     4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7,8-tetramethoxy- (9CI)
CN
        (CA INDEX NAME)
        OMe
MeO
MeO
                           OMe
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OMe

RN

CN

0

481-53-8 HCAPLUS

(CA INDEX NAME)

OMe

4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI)

RN 1721-51-3 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,5,7,8-tetramethyl-2-(4,8,12-trimethyl-3,7,11-tridecatrienyl)- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

= CMe₂

RN 6829-55-6 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2-methyl-2-(4,8,12-trimethyl-3,7,11-tridecatrienyl)- (9CI) (CA INDEX NAME)

PAGE 1-B

= CMe $_2$

RN 14101-61-2 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,7,8-trimethyl-2-[(3E,7E)-4,8,12-trimethyl-3,7,11-tridecatrienyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Page 31

Double bond geometry as shown.

RN 25612-59-3 HCAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,8-dimethyl-2-[(3E,7E)-4,8,12-trimethyl-3,7,11-tridecatrienyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

REFERENCE COUNT: 48 THERE ARE 48 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

HCAPLUS COPYRIGHT 2002 ACS L124 ANSWER 12 OF 39

2001:626002 HCAPLUS ACCESSION NUMBER:

135:185492 DOCUMENT NUMBER:

TITLE: Flavones for the treatment of COX-2 and/or

NF.kappa.B-mediated diseases Wenzel, Uwe; Daniel, Hannelore

INVENTOR(S): PATENT ASSIGNEE(S): Basf A. -G., Germany

SOURCE: Jpn. Kokai Tokkyo Koho, 13 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT:

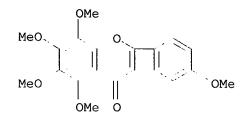
PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	JP 2001233768	A2	20010828	JP 2001-49370	20010223
	EP 1127572	A2	20010829	EP 2001-103200	20010212
	R: AT, BE,	CH, DE,	, DK, ES, FR, GE	B, GR, IT, LI, LU,	NL, SE, MC, PT,
	IE, SI,	LT, LV	, FI, RO		
	US 2001046963	A1	20011129	US 2001-782306	20010214
	CN 1318371	Α	20011024	CN 2001-116513	20010225
PRIO	RITY APPLN. INFO	.:	US	2000-185179P P	20000225
OTHE	R SOURCE(S):	MAI	RPAT 135:185492		
AB	This invention	relates	to the use of i	lavone or derivs.	. thereof for the
	treatment of di	seases r	mediated by cvc	looxygenase-2 or N	NF.kappa.B. The
				dosage forms or fo	
ΙT	481-53-8, Tange			3	
			tivity or effect	or, except advers	se): BSU
				(Food or feed use	
	-	-		udul BCEC (Hees)	

(Therapeutic use); BIOL (Biological study); USES (Uses)

(flavones for treatment of COX-2 and/or NF.kappa.B-mediated diseases)

RN 481-53-8 HCAPLUS CN 4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)



L124 ANSWER 13 OF 39 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:179725 HCAPLUS

DOCUMENT NUMBER: 132:227425

TITLE: Pharmaceuticals and foods containing flavonoids as

inhibitors of formation of matrix metalloproteinase

(MMP) and its precursor

INVENTOR(S): Yano, Masamitsu; Oqawa, Kazunori; Yoshida, Toshio;

Nezumi, Hirohisa; Nonomura, Mutsuko; Ishiwa, Atsushi; Sato, Takashi; Mitsumaki, Yoshihiro; Sashida, Yutaka;

Ito, Akira

PATENT ASSIGNEE(S): Ministry of Agriculture and Forestry National Fruits

Experiment Station, Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 12 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2000080035	A2	20000321	JP 1998-248145	19980902
TD 2010210	DΩ	20000022		

JP 3010210 B2 20000221

OTHER SOURCE(S): MARPAT 132:227425

AB The pharmaceuticals and foods are claimed. They are useful for treatment of chronic rheumatoid arthritis, osteoarthritis, tumor, arteriosclerosis, aneurysm, hepatic cirrhosis, ulcer, osteoporosis, pulmonary fibrosis, glomerular nephritis, and periodontitis. Nobiletin inhibited IL-1.alpha.-induced proMMP-9 formation as strongly as dexamethasone without affecting formation of proMMP-2.

IT 478-01-3P, Nobiletin 481-53-8P,

Tangeretin 2174-59-6P, 5-Demethylnobiletin

2306-27-6P 6601-66-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); FFD (Food or feed use); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(citrus flavonoids as inhibitors of formation of matrix

metalloproteinase for treatment of diseases)

RN 478-01-3 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7,8-tetramethoxy- (9CI) (CA INDEX NAME)

RN 481-53-8 HCAPLUS

CN 4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 2174-59-6 HCAPLUS

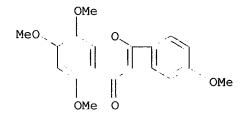
CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-6,7,8-trimethoxy-(9CI) (CA INDEX NAME)

RN 2306-27-6 HCAPLUS

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5,6,7-trimethoxy- (9CI) (CA INDEX NAME)

RN 6601-66-7 HCAPLUS

CN 4H-1-Benzopyran-4-one, 5,7,8-trimethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)



L124 ANSWER 14 OF 39 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1999:708599 HCAPLUS

DOCUMENT NUMBER: 131:317792

TITLE: Method of treatment of glutathione deficient mammals

INVENTOR(S): Keller, M. D. Robert H.; Kirchenbaum, David W.

PATENT ASSIGNEE(S): Vit-Immune, L.C., USA SOURCE: PCT Int. Appl., 26 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English .

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 9955326 A1 19991104 WO 1999-US9485 19990429

W: CA, JP

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE

US 6262019 B1 20010717 US 1999-302217 19990429 PRIORITY APPLN. INFO.: US 1998-83661P P 19980430

AB Glutathione is a tripeptide of extreme importance as a catalyst, reductant, and reactant. The disclosure is of a unique combination of nutritional supplements including N-acetylcysteine, vitamin C, L-glucosamine, N-acetyl-D-glucosamine, quercitin, sylimarin, .alpha.-lipoic acid, and high-protein, low-fat whey that are combined to support various bodily systems involved in glutathione synthesis, reutilization and storage, all intended to elevate glutathione concn. in the mammalian cell.

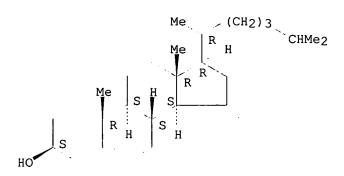
IT 57-88-5, Cholesterol, biological studies
RL: BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence); PROC (Process)

(glutathione deficiency treatment compn. and method)

RN 57-88-5 HCAPLUS

CN Cholest-5-en-3-ol (3.beta.) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L124 ANSWER 15 OF 39 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1998:31416 HCAPLUS

DOCUMENT NUMBER: 128:88155

TITLE: Method of screening foods for nutraceuticals

INVENTOR(S): Ghai, Geetha; Boyd, Charles; Csiszar, Katalin; Ho,

Chi-Tang; Rosen, Robert T.

PATENT ASSIGNEE(S): Rutgers, the State University of New Jersey, USA;

Ghai, Geetha; Boyd, Charles; Csiszar, Katalin; Ho,

Chi-Tang; Rosen, Robert T.

SOURCE: PCT Int. Appl., 70 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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KIND DATE
                                       APPLICATION NO. DATE
    PATENT NO.
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                                      WO 1997-US10368 19970620
    WO 9748823
                          19971224
                    A1
        W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH,
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            MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA,
            US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,
            GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
            GN, ML, MR, NE, SN, TD, TG
    US 5955269
                          19990921
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                                                         19960620
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    CA 2258821
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                          19971224
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                                         AU 1997-33950
    AU 9733950
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                          19980107
                                                         19970620
    EP 954609
                     Α1
                          19991110
                                         EP 1997-930022
                                                         19970620
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, FI
PRIORITY APPLN. INFO.:
                                      US 1996-670826
                                                         19960620
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AB The invention relates to an assay system for screening nutraceut

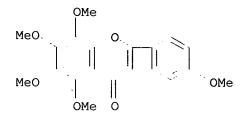
AB The invention relates to an assay system for screening nutraceuticals, i.e., foods or food substances that occur naturally, or that are produced during processing which are capable of modulating in a subject the expression of one or more genes assocd. With a disease or undesirable condition. The effect of nutraceuticals on lysyl oxidase promoter activity is shown in the figure. The nutraceuticals identified by the screening assays can be incorporated into compns. Which may be administered to a subject to treat or prevent a disease or undesirable condition, or otherwise to improve the health of the subject. The invention also provides methods for detg. the effect of a food or food substance on the expression of disease-related genes. The invention further provides methods for modifying the amt. of nutraceuticals in raw and processed foods or food substances.

IT 481-53-8, Tangeretin

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (method of screening foods for nutraceuticals)

RN 481-53-8 HCAPLUS

CN 4H-1-Benzopyran-4-one, 5,6,7,8-tetramethoxy-2-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)



L124 ANSWER 16 OF 39 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1996:161229 HCAPLUS

DOCUMENT NUMBER: 124:185594

TITLE: Pharmaceutical and cosmetic formulations containing

esculoside

INVENTOR(S): Bombardelli, Ezio; Cristoni, Aldo; Morazzoni, Paolo

PATENT ASSIGNEE(S): Indena S.p.A., Italy SOURCE: Eur. Pat. Appl., 8 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	EP 692250	A2	19960117	EP 1995-110463	19950705
	EP 692250	A3	19961023		
	EP 692250	В1	20020605		
	R: AT, BE,	CH, DE	, DK, ES,	FR, GB, GR, IE, IT, LI,	LU, MC, NL, PT, SE
	CA 2153604	AA	19960113	CA 1995-2153604	19950711
	AU 9524919	A1	19960125	AU 1995-24919	19950711
	AU 686381	В2	19980205		
	JP 08169896	A2	19960702	JP 1995-174658	19950711
)I	RITY APPLN, INFO.	:		TT 1994-MT1446 A	19940712

PRIORITY APPLN. INFO.: 11 1994-M11446 Esculoside (I) alone or in combination with adenylate cyclase stimulators, such as forskolin or Salvia miltiorrhiza diterpenes and/or with phosphodiesterase inhibitors, such as apigenin-skeleton dimeric flavones are used in topical formulations for the treatment of peripheral vasculophathies related to an impaired peripheral microcirculation, cellulitis or unesthetisms connected with a deposit of superfluous fat. For the redn. of the deposits of superfluous fat of any origin, the above mentioned products are advantageously also combined with caffeine, theophylline and derivs. thereof. Efficacy of 1.5% I in treatment of patients affected with venous insufficiency is reported. A gel contained S. miltiorrhiza ext. 0.30, I 1.50, Ginkgo biloba dimeric flavones 0.50, hydrogenated ethoxylated castor oil 1.00, propylene glycol 1.50, preservatives 0.10, hydroxyethyl cellulose 3.00, and purified water q.s. 100g.

L124 ANSWER 17 OF 39 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1996:151191 HCAPLUS

DOCUMENT NUMBER: 124:278479

TITLE: Anticholesteremic effect of flavonoid derivatives in

rat

AUTHOR(S): Nagem, Tanus Jorge; de Oliveira, Tania Toledo; da

Silva, Marilda Conceicao; Guedes de Miranda, Luiz

Carlos

CORPORATE SOURCE: Departamento de Quimica, UFV, Vicosa, 36570-000,

Brazil

SOURCE: Arq. Biol. Tecnol. (1995), Volume Date 1995, 38(3),

859-68

CODEN: ABTTAP; ISSN: 0365-0979

DOCUMENT TYPE: LANGUAGE:

Journal Portuguese

O-Me and acetyl derivs. of morin, naringenin, quercetin and rutin isolated from soya cultivar UFV-5' were prepd., identified by UV, IR, NMR and tested in rats against cholesterol. Animals that were administered quercetin derivs. and methylated rutin showed lowest concns. of lipids in the bloodstream and highest concns. of biliary salts.

IT 1245-15-4P

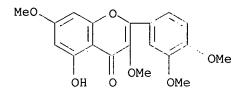
> RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological

study); PREP (Preparation); USES (Uses)

(anticholesteremic effect of flavonoid derivs. in rats)

1245-15-4 HCAPLUS RN

CN 4H-1-Benzopyran-4-one, 2-(3,4-dimethoxyphenyl)-5-hydroxy-3,7-dimethoxy-(9CI) (CA INDEX NAME)



L124 ANSWER 18 OF 39 HCAPLUS COPYRIGHT 2002 ACS

1977:577782 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

87:177782

TITLE:

Effect of flavonoids and galascorbin on some catabolic

processes and cholesterol removal during experimentally-induced hypercholesteremia

AUTHOR(S):

Kiyasheva, T. Zh.

CORPORATE SOURCE: Karagand. Med. Inst., Karaganda, USSR

SOURCE:

Fiziol. Patol. Organov Pishchevareniya (1974), 71-5. Editor(s): Dauletbakova, M. I. Karagand. Gos. Med.

Inst.: Karaganda, USSR.

CODEN: 36MOAH

DOCUMENT TYPE:

Conference

LANGUAGE:

Russian

Feeding an atherogenic diet for 30 days to rats increased the excretion of cholesterol [57-88-5] and cholic acid [81-25-4] in bile and of cholesterol and total steroids in feces. Rutin [153-18-4] (100 or 200 mg/kg), quercitin [117-39-5] (200 mg/kg), or galascorbin [8065-60-9] (100 mg/kg) given orally simultaneously with the atherogenic diet further increased cholesterol and cholic acid excretion in the bile and cholesterol and steroid excretion in the feces. Apparently, the flavonoids and galascorbin affect liver function rather than inhibit absorption of cholesterol and cholic acid by the intestine.

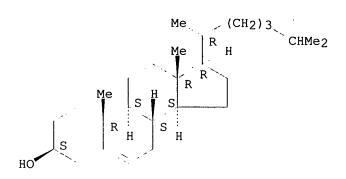
TT 57-88-5, biological studies RL: BIOL (Biological study)

(of blood serum, flavonoids and galascorbin effect on)

RN 57-88-5 HCAPLUS

CN Cholest-5-en-3-ol (3.beta.) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L124 ANSWER 19 OF 39 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1966:432882 HCAPLUS

DOCUMENT NUMBER: 65:32882
ORIGINAL REFERENCE NO.: 65:6141c-d

TITLE: Effect of a preparation of common onion skin on the

cholesterol content of blood and aorta in

experimental hypercholesterolemia in white rats

AUTHOR(S): Lisevitskaya, L. I.; Bardyukova, V. A.; Shinkarenko,

A. L.

CORPORATE SOURCE: Pharm. Inst., Pyatigorsk

SOURCE: Nauchn. Dokl. Vysshei Shkoly, Biol. Nauki (1966), (2),

78-9 Journal

DOCUMENT TYPE: Journal LANGUAGE: Russian

AB Exptl. hypercholesterolemia was induced in rats by addn. of 600 mg. cholesterol (I) and 90 mg. methylthiouracil (II/kg. body wt./day). Thus, the content of I in the blood had increased to 95 and 140 mg. % after 1 and 2 months, resp. (control animals 45-50 mg. %). Addn. of a prepn. of common onion (Allium cepa) skin (5 mg./kg. body wt./day) contg. the whole of polyphenolic compds. with 30% quercitin, decreased the concn. of I to normal after 1.5 months, though the application of I and II was continued. Detn. of I in the aorta gave the same value (106-110 mg. %) for control animals and those treated with I, II, and onion prepn., whereas in rats supplied only with I and II 186 mg. % was found.

L124 ANSWER 20 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 2002124874 EMBASE

TITLE: Ventricular remodeling by Scutellarein treatment

in spontaneously hypertensive rats. Zhou J.; Lei H.; Chen Y.; Li F.; Ma C.

CORPORATE SOURCE: J. Zhou, Department of Internal Medicine, The First

Affiliated Hospital, Chongqing Univ. of Medical Sciences,

Chongqing 400016, China

SOURCE: Chinese Medical Journal, (2002) 115/3 (375-377).

Refs: 5

ISSN: 0366-6999 CODEN: CMDJAE

COUNTRY: China

DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 018 Cardiovascular Diseases and Cardiovascular Surgery

030 Pharmacology

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

AUTHOR:

Objective. To observe reversal of ventricular remodeling by the protein kinase

09/528488 Owens Page 39

C inhibitor Scutellarein in spontaneously hypertensive rats (SHRs). Methods. Twelve SHRs were randomly divided into two groups. ***Scutellarein*** and saline (10 mg.ovrhdot.kg(-1).ovrhdot.d(-1)) were given by intraperitoneal injection to two groups of rats separately. Systolic blood pressure (SBP) and ventricular weight index (LVW/BW, RVW/BW) were measured. A polarization microscope and an image analyzer system (IAS) were used to observe changes in cardiovascular structure and to count the content of cardiac muscle interstitial collagen. Results. The pathologic changes in the left ventricle in the Scutellarein group rats (SHR(D)) improved to varying degrees, including hypertrophy of the cardiac muscle and collagen volume fraction. Conclusion. Scutellarein can reverse ventricular remodeling, improve myocardial stiffness and protect heart cardiac muscle.

CONTROLLED TERM: Medical Descriptors:

*essential hypertension: DT, drug therapy

*heart ventricle remodeling spontaneously hypertensive rat

dose response

systolic blood pressure

heart weight body weight

polarization microscope

image analysis heart muscle

heart ventricle hypertrophy

heart protection

nonhuman

rat

animal experiment animal model controlled study animal tissue

article

Drug Descriptors:

*scutellarein: DO, drug dose *scutellarein: DT, drug therapy *scutellarein: PD, pharmacology

*scutellarein: IP, intraperitoneal drug

administration

protein kinase C inhibitor: DO, drug dose protein kinase C inhibitor: DT, drug therapy protein kinase C inhibitor: PD, pharmacology

protein kinase C inhibitor: IP, intraperitoneal drug

administration sodium chloride

collagen: EC, endogenous compound

CAS REGISTRY NO.: (scutellarein) 529-53-3; (sodium chloride)

7647-14-5; (collagen) 9007-34-5

L124 ANSWER 21 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER:

TITLE:

2002180565 EMBASE

Regulation of lipoprotein metabolism in HepG2 cells by

citrus flavonoids.

AUTHOR: Kurowska E.M.; Manthey J.A.

CORPORATE SOURCE: E.M. Kurowska, KGK Synergize, Inc., 255 Queens Avenue,

London, Ont. N6A 5R8, Canada

SOURCE: Advances in Experimental Medicine and Biology, (2002) 505/-

> (173-179). Refs: 22

ISSN: 0065-2598 CODEN: AEMBAP

COUNTRY: United States

DOCUMENT TYPE: Journal; Conference Article

FILE SEGMENT: 018 Cardiovascular Diseases and Cardiovascular Surgery

Page 40

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037
                             Drug Literature Index
LANGUAGE:
                    English
CONTROLLED TERM:
                    Medical Descriptors:
                      *hypercholesterolemia
                    cell strain HepG2
                    lipoprotein metabolism
                    citrus fruit
                    orange (fruit)
                    grapefruit
                    orange juice
                    grapefruit juice
                    cholesterol metabolism
                    human
                    human cell
                    conference paper
                    priority journal
                    Drug Descriptors:
                       *flavonoid: PD, pharmacology
                    lipoprotein: EC, endogenous compound
                      isoflavone derivative: PD, pharmacology
                      genistein: PD, pharmacology
                      hesperetin: PD, pharmacology
                      naringenin: PD, pharmacology
                      hypocholesterolemic agent: PD, pharmacology
                      hesperidin: PD, pharmacology
                      aurantiin: PD, pharmacology
                      tangeretin: PD, pharmacology
                      nobiletin: PD, pharmacology
                      sinensetin: PD, pharmacology
                      scutellarein: PD, pharmacology
                      tetra o methylscutellarein: PD, pharmacology
                      antiinflammatory agent: PD, pharmacology
                      low density lipoprotein: EC, endogenous compound
                    high density lipoprotein: EC, endogenous compound
                      cholesterol: EC, endogenous compound
                      apolipoprotein B: EC, endogenous compound
                      3,5,6,7,8,3',4' heptamethoxyflavone: PD,
                    pharmacology
                      5 norsinensetin: PD, pharmacology
                      quercetin derivative: PD, pharmacology
                      quercetin 3,7,3',4' tetramethyl ether: PD,
                    pharmacology
                      quercetin 3,5,7,3',4' pentamethyl ether: PD,
                    pharmacology
                    unclassified drug
CAS REGISTRY NO.:
                    (genistein) 446-72-0; (hesperetin) 520-33-2; (naringenin)
                    480-41-1, 67604-48-2; (hesperidin) 520-26-3; (aurantiin)
                    10236-47-2, 12619-61-3, 29658-83-1, 82350-96-7; (
                    tangeretin) 481-53-8; (nobiletin
                    ) 478-01-3; (scutellarein) 529-53-3;
                    (cholesterol) 57-88-5
L124 ANSWER 22 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.
ACCESSION NUMBER:
                    2002034023 EMBASE
TITLE:
                    Upregulation of interleukin-8 expression by prostaglandin
                    D2 metabolite 15-deoxy-delta12, 14 prostaglandin J2
                    (15d-PGJ2) in human THP-1 macrophages.
AUTHOR:
                    Fu Y.; Luo N.; Lopes-Virella M.F.
CORPORATE SOURCE:
                    Y. Fu, Department of Medicine, Strom Thurmond Biomedical
                    Center, Medical University of South Carolina, 114 Doughty
                    Street, Charleston, SC 29403-5729, United States.
```

030

Pharmacology

fuv@musc.edu

SOURCE:

Atherosclerosis, (2002) 160/1 (11-20).

Refs: 39

ISSN: 0021-9150 CODEN: ATHSBL

PUBLISHER IDENT.:

S 0021-9150(01)00541-X

COUNTRY:

Ireland

DOCUMENT TYPE:

Journal; Article

FILE SEGMENT:

018 Cardiovascular Diseases and Cardiovascular Surgery

030 Pharmacology

Medical Descriptors:

037 Drug Literature Index

LANGUAGE:

English English

Science Ireland Ltd. All rights reserved.

SUMMARY LANGUAGE: ABSTRACT:

Interleukin-8 (IL-8) is one of cytokines detected at sites of inflammation and in macrophage-foam cells of atherosclerotic lesions. The expression of IL-8 gene can be induced in cholesterol loaded THP-1 macrophages by oxidized low density lipoprotein. We report for the first time that the expression of human IL-8 gene in THP-1 macrophages is upregulated in a time- and concentration-dependent manner by prostaglandin D2 metabolite 15-deoxy-delta12, 14 prostaglandin J2 (15d-PGJ2), which is a natural ligand for activation of peroxisome proliferator-activated receptor-gamma transcription factor. Studies to identify the signal transduction pathways involved showed that IL-8 upregulation-mediated by 15d-PGJ2 was markedly inhibited when the THP-1 macrophages were incubated with a highly selective and cell-permeable inhibitor of the mitogen-activated protein kinase 02(MAPK) signaling pathway, 2'-amino-3'-methoxyflavone (PD98059). This inhibition was concentration-dependent, suggesting that 15d-PGJ2 regulates the expression of IL-8 gene in THP-1 macrophages through a MAPK signaling pathway. In contrast, THP-1 macrophages when treated with pyrrolidine dithiocarbamate, an anti-oxidant and the selective inhibitor for nuclear factor .kappa.B, showed an enhanced 15d-PGJ2-mediated upregulation of IL-8 gene expression. The data presented in this report may contribute to unravel some of the mechanisms behind the inflammatory component of atherosclerosis. . COPYRGT. 2002 Elsevier

CONTROLLED TERM:

*atherosclerosis *signal transduction metabolite protein expression regulatory mechanism macrophage inflammation foam cell peroxisome incubation time concentration response gene expression human controlled study human cell article priority journal Drug Descriptors: *interleukin 8: EC, endogenous compound *prostaglandin D2: EC, endogenous compound *delta12 prostaglandin J2: EC, endogenous compound *2 (2 amino 3 methoxyphenyl)chromone: PD, pharmacology *pyrrolidine dithiocarbamate: PD, pharmacology

transcription factor: EC, endogenous compound

cholesterol: EC, endogenous compound

mitogen activated protein kinase

immunoglobulin enhancer binding protein: EC, endogenous

compound

CAS REGISTRY NO.: (interleukin 8) 114308-91-7; (prostaglandin D2) 41598-07-6;

(delta12 prostaglandin J2) 87893-54-7; (2 (2 amino 3

methoxyphenyl)chromone) 167869-21-8; (cholesterol) 57-88-5;

(mitogen activated protein kinase) 142243-02-5

CHEMICAL NAME:

(1) Pd 98059

COMPANY NAME:

(1) Calbiochem (United States)

L124 ANSWER 23 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

2002180550 EMBASE ACCESSION NUMBER:

TITLE: Flavonoids in cell function.

AUTHOR: Manthey J.A.; Buslig B.S.; Baker M.E.

CORPORATE SOURCE: J.A. Manthey, U.S. Department of Agriculture, Citrus and

Subtropical Products Lab., 600 Avenue S, NW, Winter Haven,

FL 33881, United States

SOURCE: Advances in Experimental Medicine and Biology, (2002) 505/-

(1-7).

Refs: 25

ISSN: 0065-2598 CODEN: AEMBAP

COUNTRY:

United States

DOCUMENT TYPE: Journal; Conference Article FILE SEGMENT: 029 Clinical Biochemistry

030 Pharmacology

037 Drug Literature Index

LANGUAGE: English

CONTROLLED TERM:

Medical Descriptors:

*cell function microorganism higher plant pollen germination

bell pepper

citrus fruit

fruit vegetable fruit juice

estrogen activity

antiinflammatory activity antioxidant activity

ischemic heart disease

drug activity drug isolation phytochemistry

human nonhuman

conference paper priority journal Drug Descriptors:

*flavonoid: PD, pharmacology *isoflavonoid: PD, pharmacology phenol derivative: PD, pharmacology polyphenol derivative: PD, pharmacology flavonol derivative: PD, pharmacology

galactosyltransferase: EC, endogenous compound

flavonol 3 o galactosyltransferase: EC, endogenous compound

indoleacetic acid: EC, endogenous compound

alpha tocopherol

antioxidant: PD, pharmacology

antithrombocytic agent: PD, pharmacology

catechin: PD, pharmacology hesperidin: PD, pharmacology flavone derivative: PD, pharmacology methoxyflavone: PD, pharmacology tangeretin: PD, pharmacology uvomorulin: EC, endogenous compound catenin: EC, endogenous compound

interleukin 2 receptor: EC, endogenous compound

tamoxifen

immunosuppressive agent

xanthine oxidase: EC, endogenous compound xanthine dehydrogenase: EC, endogenous compound

steroid receptor: EC, endogenous compound

estrogen: EC, endogenous compound phytoestrogen: PD, pharmacology

testosterone 17beta dehydrogenase: EC, endogenous compound

adenosine receptor: EC, endogenous compound

adenosine: EC, endogenous compound

unindexed drug unclassified drug

CAS REGISTRY NO.:

(galactosyltransferase) 9031-68-9; (indoleacetic acid) 32536-43-9, 87-51-4; (alpha tocopherol) 1406-18-4, 1406-70-8, 52225-20-4, 58-95-7, 59-02-9; (catechin) 13392-26-2, 154-23-4; (hesperidin) 520-26-3; (

tangeretin) 481-53-8; (uvomorulin)

112956-45-3; (tamoxifen) 10540-29-1; (xanthine oxidase)

9002-17-9; (xanthine dehydrogenase) 9054-84-6;

(testosterone 17beta dehydrogenase) 9028-62-0; (adenosine)

58-61-7

L124 ANSWER 24 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER:

2000146748 EMBASE

TITLE:

Effects of Scutellarein on diabetic rat aorta.

AUTHOR:

Zhu B.-H.; Guan Y.-Y.; He H.; Lin M.-J.

CORPORATE SOURCE: Dr. B.-H. Zhu, Department of Pharmacology, Sun Yat-Sen

Univ. of Med. Sciences, Guangzhou 510089, China.

sszhu@gzsums.edu.cn

SOURCE:

Acta Pharmacologica Sinica, (2000) 21/4 (353-356).

Refs: 14

ISSN: 0253-9756 CODEN: CYLPDN

COUNTRY:

China

DOCUMENT TYPE: FILE SEGMENT:

Journal; Article 003 Endocrinology

018 Cardiovascular Diseases and Cardiovascular Surgery

030 Pharmacology

037 Drug Literature Index

LANGUAGE:

English

SUMMARY LANGUAGE:

English; Chinese

ABSTRACT:

AIM: To study the effect of **Scutellarein** (Scu) on the diabetic rat aorta. METHODS: Contractile responses to phenylepherine and endothelium-dependent relaxation responses to acetylcholine (ACh) in rat aorta were investigated after streptozocin-induced 6-wk diabetes, Scu-treated streptozocin-induced diabetes, and in age-matched control in vitro. RESULTS: 1) Endothelium-dependent relaxation to ACh in diabetic rats was decreased (P < 0.01) compared with age-matched control. 2) Contractile responses to phenylepherine were increased (P < 0.01) in diabetic rats. 3) The dietary supplement of 0.5 % Scu starting from 1-wk diabetes induction prevented endothelial dysfunction (P < 0.01), but the contractile responses to phenylepherine were further increased. CONCLUSION: Scu prevented vascular endothelial dysfunction in diabetic rats, and also potentiated the contraction induced by phenylepherine.

CONTROLLED TERM:

Medical Descriptors:

*thoracic aorta

*streptozocin diabetes *diabetic angiopathy: DT, drug therapy *diabetic angiopathy: PC, prevention *endothelium injury: DT, drug therapy *endothelium injury: PC, prevention vascular endothelium vasoconstriction vasodilatation treatment outcome nonhuman male rat animal experiment animal model controlled study animal tissue article Drug Descriptors: *scutellarein: IT, drug interaction *scutellarein: DT, drug therapy *scutellarein: PD, pharmacology *scutellarein: PO, oral drug administration phenylephrine: IT, drug interaction phenylephrine: PD, pharmacology acetylcholine streptozocin (scutellarein) 529-53-3; (phenylephrine) 532-38-7, 59-42-7, 61-76-7; (acetylcholine) 51-84-3, 60-31-1, 66-23-9; (streptozocin) 18883-66-4 Otsuka; Sigma L124 ANSWER 25 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V. 2000037681 EMBASE Hepatocellular carcinoma. Badvie S. S. Badvie, Surgical Unit, St. Thomas's Hospital, Lambeth Palace Road, London SE1, United Kingdom Postgraduate Medical Journal, (2000) 76/891 (4-11). Refs: 108 ISSN: 0032-5473 CODEN: PGMJAO United Kingdom Journal; Article 016 Cancer 048 Gastroenterology 026 Immunology, Serology and Transplantation 037 Drug Literature Index 014 Radiology English English

LANGUAGE: SUMMARY LANGUAGE:

ABSTRACT:

CAS REGISTRY NO.:

ACCESSION NUMBER:

CORPORATE SOURCE:

COMPANY NAME:

TITLE:

AUTHOR:

SOURCE:

COUNTRY:

DOCUMENT TYPE:

FILE SEGMENT:

Primary hepatocellular carcinoma is one of the 10 most common tumours, and the most common primary liver malignancy, in the world. In the majority of cases, it occurs against a background of hepatitis B or C viral infection and/or liver cirrhosis, and is associated with a dismal prognosis of a few months. Current treatments in routine clinical practice are surgical resection and liver transplantation, but these therapies are applicable to only a small proportion of patients and prolongation of survival is restricted. Other treatment options include intra-arterial chemotherapy, transcatheter arterial chemoembolisation, percutaneous ethanol injection, cryotherapy, thermotherapy, proton therapy, or a wide range of their possible combinations. The current lack of definitive data, however, limits the use of these therapies. Another option is gene therapy, which although in its infancy at the present time, may have a significant role to play in the future management of hepatocellular carcinoma.

CONTROLLED TERM: Medical Descriptors: *liver cell carcinoma: DI, diagnosis *liver cell carcinoma: SU, surgery *liver cell carcinoma: DT, drug therapy *liver cell carcinoma: TH, therapy *liver cell carcinoma: RT, radiotherapy *liver cell carcinoma: PC, prevention human major clinical study controlled study randomized controlled trial hepatitis C: ET, etiology hepatitis C: PC, prevention hepatitis B: ET, etiology hepatitis B: PC, prevention liver cirrhosis: ET, etiology liver transplantation artificial embolism cryotherapy clinical trial multidrug resistance fast proton radiation hyperthermic therapy pancreatitis: CO, complication peptic ulcer: CO, complication necrosis: CO, complication liver failure: CO, complication liver abscess: CO, complication arteritis: CO, complication gallbladder disease: CO, complication immunotherapy herbal medicine article Drug Descriptors: *epirubicin: DT, drug therapy *epirubicin: CT, clinical trial *mitoxantrone: DT, drug therapy *mitoxantrone: CT, clinical trial *platinum complex: DT, drug therapy *platinum complex: CT, clinical trial *amsacrine: DT, drug therapy *amsacrine: CT, clinical trial *fludarabine: DT, drug therapy *fludarabine: CT, clinical trial *vinblastine: DT, drug therapy *vinblastine: CT, clinical trial *zidovudine: DT, drug therapy *zidovudine: CT, clinical trial *doxifluridine: DT, drug therapy *doxifluridine: CT, clinical trial *fluorouracil: DT, drug therapy *fluorouracil: IA, intraarterial drug administration *fluorouracil: CT, clinical trial *anthracycline: DT, drug therapy *anthracycline: IA, intraarterial drug administration *anthracycline: CT, clinical trial *floxuridine: DT, drug therapy *floxuridine: IA, intraarterial drug administration *floxuridine: CB, drug combination *floxuridine: CT, clinical trial *folinic acid: DT, drug therapy

*folinic acid: IA, intraarterial drug administration

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*folinic acid: CB, drug combination
                    *folinic acid: CT, clinical trial
                      *cisplatin: DT, drug therapy
                    *cisplatin: IA, intraarterial drug administration
                    *cisplatin: CB, drug combination
                    *cisplatin: CT, clinical trial
                      *doxorubicin: DT, drug therapy
                    *doxorubicin: IA, intraarterial drug administration
                    *doxorubicin: CB, drug combination
                    *doxorubicin: CT, clinical trial
                      *mitomycin: DT, drug therapy
                    *mitomycin: IA, intraarterial drug administration
                    *mitomycin: CT, clinical trial
                    *mitoxantrone: IA, intraarterial drug administration
                    *gelatin sponge
                      *alcohol: DT, drug therapy
                      *tamoxifen: DT, drug therapy
                    *tamoxifen: CT, clinical trial
                      *flutamide: DT, drug therapy
                    *flutamide: CT, clinical trial
                      *ketoconazole: DT, drug therapy
                    *ketoconazole: CT, clinical trial
                      *buserelin: DT, drug therapy
                    *buserelin: CT, clinical trial
                      *retinoic acid: DT, drug therapy
                    *retinoic acid: CT, clinical trial
                      *octreotide: DT, drug therapy
                    *octreotide: CT, clinical trial
                      *inchinko to: DT, drug therapy
                      *flavinoid quercitin: DT, drug therapy
                      *hepatitis B vaccine: DT, drug therapy
CAS REGISTRY NO.:
                    (epirubicin) 56390-09-1, 56420-45-2; (mitoxantrone)
                    65271-80-9, 70476-82-3; (amsacrine) 51264-14-3, 54301-15-4;
                    (fludarabine) 21679-14-1; (vinblastine) 865-21-4;
                    (zidovudine) 30516-87-1; (doxifluridine) 3094-09-5;
                    (fluorouracil) 51-21-8; (floxuridine) 50-91-9; (folinic
                    acid) 58-05-9, 68538-85-2; (cisplatin) 15663-27-1,
                    26035-31-4, 96081-74-2; (doxorubicin) 23214-92-8,
                    25316-40-9; (mitomycin) 1404-00-8; (mitoxantrone)
                    65271-80-9, 70476-82-3; (alcohol) 64-17-5; (tamoxifen)
                    10540-29-1; (flutamide) 13311-84-7; (ketoconazole)
                    65277-42-1; (buserelin) 57982-77-1; (retinoic acid)
                    302-79-4; (octreotide) 83150-76-9
L124 ANSWER 26 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.
ACCESSION NUMBER:
                    2000382783 EMBASE
                    The oestrogen receptor and its selective modulators in
TITLE:
                    gynaecological and breast cancer.
AUTHOR:
                    Vergote I.; Neven P.; Van Dam P.; Serreyn R.; De Prins F.;
                    De Sutter P.; Albertyn G.
CORPORATE SOURCE:
                    I. Vergote, Gynaecological Oncology, University Hospitals
                    Leuven, Herestraat 49, B-3000 Leuven, Belgium.
                    ignace.vergote@uz.kuleuven.ac.be
SOURCE:
                    European Journal of Cancer, (2000) 36/SUPPL. 4 (S1-S9).
                    Refs: 70
                    ISSN: 0959-8049 CODEN: EJCAEL
                    S 0959-8049(00)00203-3
PUBLISHER IDENT.:
COUNTRY:
                    United Kingdom
DOCUMENT TYPE:
                    Journal; General Review
FILE SEGMENT:
                    010
                            Obstetrics and Gynecology
                    016
                            Cancer
                    030
                             Pharmacology
                    037
                             Drug Literature Index
```

038 Adverse Reactions Titles LANGUAGE: English CONTROLLED TERM: Medical Descriptors: *gynecologic cancer: DT, drug therapy *breast carcinoma: DT, drug therapy *drug mechanism hormonal therapy protein domain menopause drug receptor binding uterus carcinoma hormone responsive element cancer survival amino terminal sequence endometrium carcinoma: SI, side effect vagina bleeding: SI, side effect thromboembolism: SI, side effect human major clinical study human tissue review priority journal Drug Descriptors: *estrogen receptor *selective estrogen receptor modulator: DT, drug therapy *selective estrogen receptor modulator: PD, pharmacology tamoxifen: AE, adverse drug reaction tamoxifen: CM, drug comparison tamoxifen: PD, pharmacology tamoxifen: PO, oral drug administration estrogen receptor alpha estrogen receptor beta transcription factor toremifene: AE, adverse drug reaction toremifene: CM, drug comparison toremifene: PD, pharmacology idoxifene: PD, pharmacology benzothiophene derivative naphthalene derivative benzopyran derivative nafoxidine: AE, adverse drug reaction trioxifene: AE, adverse drug reaction zindoxifene: AE, adverse drug reaction arzoxifene: CM, drug comparison arzoxifene: PD, pharmacology raloxifene: CM, drug comparison raloxifene: PD, pharmacology 7alpha [9 (4,4,5,5,5 pentafluoropentylsulfinyl)nonyl]e stra 1,3,5(10) triene 3,17beta diol: PD, pharmacology anastrozole: AE, adverse drug reaction anastrozole: CM, drug comparison anastrozole: PD, pharmacology 11 [4 [[5 [(4,4,5,5,5 pentafluoropentyl)sulfonyl]penty l]oxy]phenyl]estradiol: PD, pharmacology medroxyprogesterone acetate: CM, drug comparison medroxyprogesterone acetate: PD, pharmacology medroxyprogesterone acetate: PO, oral drug administration isoflavone phytoestrogen

tangeretin: DV, drug development

aromatase inhibitor
aminoglutethimide

letrozole: CM, drug comparison
letrozole: PD, pharmacology

exemestane

megestrol acetate: CM, drug comparison
megestrol acetate: PD, pharmacology

fadrozole vorozole unindexed drug

CAS REGISTRY NO.: (tamoxifen) 10540-29-1; (toremifene) 89778-26-7;

(idoxifene) 116057-75-1; (nafoxidine) 1845-11-0, 1847-63-8;

(trioxifene) 63619-84-1; (zindoxifene) 86111-26-4; (arzoxifene) 182133-25-1, 182133-27-3; (raloxifene)

82640-04-8, 84449-90-1; (7alpha [9 (4,4,5,5,5) pentafluoropentylsulfinyl)nonyl]estra 1,3,5(10) triene 3,17beta diol) 129453-61-8; (anastrozole) 120511-73-1; (11 [4 [[5 [(4,4,5,5,5 pentafluoropentyl)sulfonyl]pentyl]oxy]ph enyl]estradiol) 151555-47-4; (medroxyprogesterone acetate)

71-58-9; (isoflavone) 574-12-9; (tangeretin)

481-53-8; (aminoglutethimide) 125-84-8; (letrozole)

112809-51-5; (exemestane) 107868-30-4; (megestrol acetate) 595-33-5; (fadrozole) 102676-31-3; (vorozole) 118949-22-7,

129731-10-8

CHEMICAL NAME: Ru 58668; Ici 182780; Ly 353381

L124 ANSWER 27 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 2000055580 EMBASE

TITLE: Flavonoids - A review of biological activities.

AUTHOR: Jaggi R.K.; Kapoor S.

CORPORATE SOURCE: R.K. Jaggi, Univ. Inst. of Pharmaceutical Sci., Panjab

University, Chandigarh 160014, India Indian Drugs, (1999) 36/11 (668-678).

Refs: 153

ISSN: 0019-462X CODEN: INDRBA

COUNTRY: India

DOCUMENT TYPE: Journal; General Review

FILE SEGMENT: 037 Drug Literature Index

030 Pharmacology

038 Adverse Reactions Titles

LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

SOURCE:

Flavonoids - a group of phenolic derivatives with diverse chemical structure, are widely distributed in plants. Flavonoids have a variety of biological activities and recently this group of natural products has gained much interest as bioactive compounds. This review gives an account of various biological activities of flavonoids.

CONTROLLED TERM: Medical Descriptors:

*drug activity

human nonhuman plant

vascular disease: DT, drug therapy vascular disease: PC, prevention

antineoplastic activity antiinflammatory activity

antiviral activity cosmetic industry liver protection drug inhibition

ulcer: DT, drug therapy

Page 49

ulcer: SI, side effect cancer: DT, drug therapy inflammatory disease: DT, drug therapy liver disease: DT, drug therapy antimicrobial activity infection: DT, drug therapy antioxidant activity cardiotoxicity review Drug Descriptors: *flavonoid: DT, drug therapy *flavonoid: PD, pharmacology *flavonoid: PO, oral drug administration *flavonoid: IV, intravenous drug administration flavone: DT, drug therapy hesperidin: DT, drug therapy hesperidin: PD, pharmacology quercitrin: DT, drug therapy quercitrin: PD, pharmacology progesterone: DT, drug therapy progesterone: PD, pharmacology naringenin: DT, drug therapy naringenin: PD, pharmacology kaempferol: DT, drug therapy kaempferol: PD, pharmacology silymarin: DT, drug therapy silymarin: PD, pharmacology gossypetin: DT, drug therapy gossypetin: PD, pharmacology acetylsalicylic acid: AE, adverse drug reaction ascorbic acid: PD, pharmacology hinokiflavone: DT, drug therapy hinokiflavone: PD, pharmacology monoxerutin: DT, drug therapy monoxerutin: PD, pharmacology troxerutin: DT, drug therapy troxerutin: PD, pharmacology quercetin 3 methyl ether: DT, drug therapy quercetin 3 methyl ether: PD, pharmacology fisetin: DT, drug therapy fisetin: PD, pharmacology taxifolin: DT, drug therapy taxifolin: PD, pharmacology tangeretin: DT, drug therapy tangeretin: PD, pharmacology luteolin: DT, drug therapy luteolin: PD, pharmacology hesperetin: DT, drug therapy hesperetin: PD, pharmacology apigenin: DT, drug therapy apigenin: PD, pharmacology acacetin: DT, drug therapy acacetin: PD, pharmacology acacetin: PO, oral drug administration rutoside derivative: DT, drug therapy rutoside derivative: PD, pharmacology esculetin: DT, drug therapy esculetin: PD, pharmacology s adenosylmethionine: DT, drug therapy s adenosylmethionine: PD, pharmacology doxorubicin: TO, drug toxicity apiin: DT, drug therapy apiin: PD, pharmacology

papaverine: DT, drug therapy papaverine: PD, pharmacology genistein: DT, drug therapy genistein: PD, pharmacology

unindexed drug chromocor

flavo ce

CAS REGISTRY NO.: (flavone) 525-82-6; (hesperidin) 520-26-3; (quercitrin)

522-12-3; (progesterone) 57-83-0; (naringenin) 480-41-1, 67604-48-2; (kaempferol) 520-18-3; (silymarin) 65666-07-1; (gossypetin) 489-35-0; (acetylsalicylic acid) 493-53-8, 50-78-2, 53663-74-4, 53664-49-6, 63781-77-1; (ascorbic acid) 134-03-2, 15421-15-5, 50-81-7; (hinokiflavone) 19202-36-9; (monoxerutin) 55965-63-4; (troxerutin) 7085-55-4, 84932-19-4; (quercetin 3 methyl ether) 1486-70-0; (fisetin) 528-48-3; (taxifolin) 480-18-2; (

tangeretin) 481-53-8; (luteolin)

491-70-3; (hesperetin) 520-33-2; (apigenin) 520-36-5;

(acacetin) 480-44-4; (esculetin) 305-01-1; (s

adenosylmethionine) 29908-03-0, 485-80-3; (doxorubicin) 23214-92-8, 25316-40-9; (apiin) 26544-34-3; (papaverine)

58-74-2, 61-25-6; (genistein) 446-72-0

CHEMICAL NAME: Chromocor; Flavo ce

L124 ANSWER 28 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 1999042736 EMBASE

TITLE: Influence of the antioxidant quercetin in vivo on the level

of nitric oxide determined by electron paramagnetic resonance in rat brain during global ischemia and

reperfusion.

AUTHOR: Shutenko Z.; Henry Y.; Pinard E.; Seylaz J.; Potier P.;

Berthet F.; Girard P.; Sercombe R.

CORPORATE SOURCE: Dr. R. Sercombe, UPR 646 CNRS, Universite Paris VII, 10

Avenue de Verdun, 75010 Paris, France.

sercombe@ext.jussieu.fr

SOURCE: Biochemical Pharmacology, (1999) 57/2 (199-208).

Refs: 66

ISSN: 0006-2952 CODEN: BCPCA6

PUBLISHER IDENT.: S 0006-2952(98)00296-2

COUNTRY: United States

DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 029 Clinical Biochemistry

030 Pharmacology

037 Drug Literature Index 008 Neurology and Neurosurgery

LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

We characterized the changes in nitric oxide (NO) levels in the brain during global forebrain ischemia and reperfusion and tested the ability of the natural flavonoid, quercetin, and a synthetic flavonoid, FB277, to increase the amount of available NO by elimination of the superoxide radicals produced during reperfusion. In Sprague-Dawley rats, we used a four-vessel occlusion model of forebrain ischemia (15 min) and reperfusion (30 min). Brain NO was measured on samples of cerebral cortex and cerebellum ex vivo by electron paramagnetic resonance (EPR) spectroscopy. The spin trap used was diethyldithiocarbamate sodium salt (DETC) associated with ferrous citrate. The complex Fe(DETC)2NO was detected at 77 K as a triplet signal at g = 2.035. Groups of animals were treated with quercetin or FB277 (3-morpholinomethyl-3',4',5,7-***tetramethoxyflavone***) or polyethylene glycol-conjugated superoxide dismutase (PEG-SOD). In control (intact anesthetized animals), the signal was about 3 times greater in the cortex than in the cerebellum. During ischemia, the signal rose to 110% in cortex (NS) and 283% in cerebellum (P < 0.05). In

reperfusion, it fell again to 91% of control in cerebellum (NS) and 35% in cortex (P < 0.05). Treatment by quercetin (5 mg/kg i.v.) of intact and ischemia-reperfusion groups did not significantly change the signal amplitude in the cerebellum, but did double it in the cortex (to 76% of control) for the ischemia-reperfusion group (P < 0.05). In contrast, FB277 (3.75 mg/kg i.v.) did not increase the signal in the cortex during ischemia-reperfusion, but did do so in the cerebellum (to 152% of control, P < 0.05). The results obtained for PEG-SOD (10,000 U/kg i.v.) were similar to those for FB277. In separate in vitro measurements, we found that quercetin but not FB277 efficiently scavenged superoxide. We hypothesize that quercetin but not FB277 scavenged superoxide anions released in the cortex during reperfusion, thus diminishing the amount of NO removed by the formation of peroxynitrite. The lack of effect of PEG-SOD may be related to the need for chronic treatment to obtain protection. Copyright (C) 1999 Elsevier Science Inc.

CONTROLLED TERM: Medical Descriptors:

*electron spin resonance

*brain ischemia: DT, drug therapy

*reperfusion nonhuman male rat

animal experiment
animal model
controlled study

intravenous drug administration
intraperitoneal drug administration

article

priority journal
Drug Descriptors:

*nitric oxide: EC, endogenous compound

*quercetin: PD, pharmacology
*quercetin: DT, drug therapy
*quercetin: CM, drug comparison
*flavonoid: PD, pharmacology
*flavonoid: DT, drug therapy
*flavonoid: DV, drug development
*flavonoid: CM, drug comparison

*scavenger: PD, pharmacology

*superoxide dismutase macrogol: PD, pharmacology *superoxide dismutase macrogol: DT, drug therapy *superoxide dismutase macrogol: CM, drug compàrison

antioxidant: PD, pharmacology antioxidant: DT, drug therapy antioxidant: CM, drug comparison

superoxide: EC, endogenous compound CAS REGISTRY NO.: (nitric oxide) 10102-43-9; (quercetin) 117-39-5;

(superoxide) 11062-77-4

COMPANY NAME: Sigma

L124 ANSWER 29 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 97267059 EMBASE

DOCUMENT NUMBER: 1997267059

TITLE: Trypanocidal flavonoids from Trixis vauthieri.

AUTHOR: Ribeiro A.; Pilo-Veloso D.; Romanha A.J.; Zani C.L. CORPORATE SOURCE: C.L. Zani, Departamento de Quimica-ICEx-UFMG, CEP

31270-901, Av. Antonio Carlos 6627, CEP 31270-901 Belo Horizonte, MG, Brazil. zani@dcc001.ciet.fiocruz.br

SOURCE: Journal of Natural Products, (1997) 60/8 (836-838).

Refs: 29

ISSN: 0163-3864 CODEN: JNPRDF

COUNTRY: United States
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 004 Microbiology

030 Pharmacology

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

The crude extract of Trixis vauthieri (Asteraceae) was active against the trypomastigote forms of Trypanosoma cruzi, the protozoan that causes Chagas' disease. Bioassay-guided fractionation of this extract afforded the trypanocidal flavonoids 5,4'-dihydroxy-7-methaxyflavanone (1) and 5,4'-dihydroxy-3,6,7-trimethoxyflavanone (2) besides the inactive flavonoids 3,5,4'- trihydroxy-7-methoxyflavanone (3) and 5,4'-dihydroxy-3,6,7,8-tetramethoxy flavone (4). The trypanocidal activity of 1 and 2 and the presence of compounds 2 and 4 in Trixis vauthieri are reported here for the first time.

CONTROLLED TERM: Medical Descriptors:

*chagas disease: DT, drug therapy *chagas disease: ET, etiology

*trypanosoma cruzi animal experiment animal model article

blood transfusion controlled study disease carrier drug screening hemiptera mouse nonhuman

plant leaf
trypomastigote
Drug Descriptors:

*5,4' dihydroxy 3,6,7 trimethoxyflavone: PD, pharmacology

*5,4' dihydroxy 3,6,7 trimethoxyflavone: DT, drug therapy

*5,4' dihydroxy 3,6,7 trimethoxyflavone: DV, drug development

*5,4' dihydroxy 7 methoxyflavanone: DV, drug development *5,4' dihydroxy 7 methoxyflavanone: PD, pharmacology

*5,4' dihydroxy 7 methoxyflavanone: DT, drug therapy

*antitrypanosomal agent: DV, drug development
 *antitrypanosomal agent: DT, drug therapy
 *antitrypanosomal agent: PD, pharmacology

*flavonoid: PD, pharmacology *flavonoid: DT, drug therapy *flavonoid: DV, drug development *plant extract: PD, pharmacology *plant extract: DT, drug therapy

*plant extract: DV, drug development benznidazole: DT, drug therapy

crystal violet

nifurtimox: DT, drug therapy

unclassified drug

CAS REGISTRY NO.: (benznidazole) 22994-85-0; (crystal violet) 467-63-0, 548-62-9; (nifurtimox) 23256-30-6

L124 ANSWER 30 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 93273634 EMBASE

DOCUMENT NUMBER: 1993273634

TITLE: Endothelium-dependent vasorelaxing activity of wine and

other grape products.

AUTHOR: Fitzpatrick D.F.; Hirschfield S.L.; Coffey R.G.

CORPORATE SOURCE: Dept. of Pharmacology/Therapeutics, Univ. of South Florida

Coll. of Med., MDC Box 9, 12901 Bruce B. Downs Blvd., Tampa,

FL 33612-4799, United States

SOURCE: American Journal of Physiology - Heart and Circulatory

Physiology, (1993) 265/2 34-2 (H774-H778).

ISSN: 0002-9513 CODEN: AJPPDI

COUNTRY: United States
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 018 Cardiovascular Diseases and Cardiovascular Surgery

030 Pharmacology

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

Current interest in the presumed benefits of wine in protecting against coronary heart disease prompted us to investigate possible effects of various grape products on vascular function in vitro. Certain wines, grape juices, and grape skin extracts relaxed precontracted smooth muscle of intact rat aortic rings but had no effect on aortas in which the endothelium had been removed. ***Quercitin*** and tannic acid, compounds known to be present in grape skins, also produced endothelium-dependent relaxation; two other grape skin compounds, resveratrol and malvidin, did not relax the rings. Phenylephrineinduced contractions were attenuated by prior exposure of aortic rings to grape skin extracts. The extracts also increased guanosine 3',5'-cyclic monophosphate (cGMP) levels in intact vascular tissue, and both relaxation and the increase in cGMP were reversed by N(G)-monomethyl-L-arginine and N(G)-nitro-L-arginine, competitive inhibitors of the synthesis of the endothelium-derived relaxing factor, nitric oxide (NO). The vasorelaxation induced by grape products therefore appears to be mediated by the NO-cGMP pathway. If such responses occur in vivo, they could conceivably help to maintain a patent coronary artery and thereby possibly contribute to a reduced incidence of coronary heart disease.

CONTROLLED TERM: Medical Descriptors:

*ischemic heart disease: PC, prevention

*vascular endothelium

*vasodilatation animal tissue

article

concentration response

controlled study

fruit male nonhuman

priority journal

rat

smooth muscle

wine

Drug Descriptors:
*plant extract
arginine derivative

cyclic gmp: EC, endogenous compound

nitric oxide phenylephrine quercetin tannin

CAS REGISTRY NO.: (cyclic gmp) 7665-99-8; (nitric oxide) 10102-43-9;

(phenylephrine) 532-38-7, 59-42-7, 61-76-7; (quercetin)

117-39-5; (tannin) 1401-55-4

L124 ANSWER 31 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 94094940 EMBASE

DOCUMENT NUMBER: 1994094940

TITLE: Evaluation of some flavonoids as potential bradykinin

antagonists.

AUTHOR: Hye Sook Yun-Choi; Sung Hyun Chung; Young Joo Kim CORPORATE SOURCE: Natural Products Research Institute, Seoul National

University, Seoul 110-460, Korea, Republic of

SOURCE: Archives of Pharmacal Research, (1993) 16/4 (283-288).

ISSN: 0253-6269 CODEN: APHRDQ

COUNTRY: Korea, Republic of DOCUMENT TYPE: Journal; Article FILE SEGMENT: 030 Pharmacology

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

Fourteen flavonoids were evaluated for their effects as potential bradykinin (BK) antagonists. The compounds were evaluated in several in vitro and in vivo (oral administration) systems; inhibition of BK induced contractions in isolated rat ileum and uterus, antagonistic effects of BK induced plasma extravasation, reduction of acetic acid induced writhing nociception and protection from endotoxic shock. Skullcapflavone II (3), baicalein (5), 5***methoxyflavone*** (11), 6-methoxyflavone (12) and 2'***methoxyflavone*** (14) showed effects in all the tests although the order

methoxyflavone (14) showed effects in all the tests although the order of potency were somewhat varied.

CONTROLLED TERM:

Medical Descriptors:

*analgesia

*extravasation

*shock: DT, drug therapy *shock: PC, prevention *smooth muscle contraction

animal experiment
animal model
animal tissue

article

drug antagonism

ileum male mouse nonhuman

oral drug administration

rat uterus

Drug Descriptors:

*baicalein: PD, pharmacology
*baicalein: CM, drug comparison
*baicalein: DT, drug therapy
*baicalein: IT, drug interaction
*baicalein: CB, drug combination
*flavonoid: DT, drug therapy
*flavonoid: CB, drug combination
*flavonoid: CM, drug comparison
*flavonoid: PD, pharmacology

*flavonoid: PD, pharmacology
*flavonoid: IT, drug interaction

2' methoxyflavone: PD, pharmacology 2' methoxyflavone: CB, drug combination 2' methoxyflavone: CM, drug comparison 2' methoxyflavone: IT, drug interaction

2' methoxyflavone: DT, drug therapy

2',5,6' trihydroxy 7,8 dimethoxyflavone: PD, pharmacology

2',5,6' trihydroxy 7,8 dimethoxyflavone: DT, drug

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therapy
  2',5,6' trihydroxy 7,8 dimethoxyflavone: IT, drug
interaction
  2',5,6' trihydroxy 7,8 dimethoxyflavone: CM, drug
comparison
  2',5,6' trihydroxy 7,8 dimethoxyflavone: CB, drug
combination
3 hydroxyflavone: CB, drug combination
3 hydroxyflavone: CM, drug comparison
  3 hydroxyflavone: PD, pharmacology
  3 hydroxyflavone: DT, drug therapy
3 hydroxyflavone: IT, drug interaction
  5 methoxyflavone: CB, drug combination
  5 methoxyflavone: CM, drug comparison
  5 methoxyflavone: DT, drug therapy
  5 methoxyflavone: PD, pharmacology
  5 methoxyflavone: IT, drug interaction
  6 methoxyflavone: PD, pharmacology
  6 methoxyflavone: DT, drug therapy
  6 methoxyflavone: IT, drug interaction
  6 methoxyflavone: CB, drug combination
  6 methoxyflavone: CM, drug comparison
apigenin: CB, drug combination
  apigenin: PD, pharmacology
apigenin: IT, drug interaction
apigenin: CM, drug comparison
  apigenin: DT, drug therapy
bradykinin: IT, drug interaction
bradykinin: TO, drug toxicity
  bradykinin: PD, pharmacology
bradykinin: CB, drug combination
chrysin dimethyl ether: CB, drug combination
  chrysin dimethyl ether: DT, drug therapy
  chrysin dimethyl ether: PD, pharmacology
chrysin dimethyl ether: IT, drug interaction
chrysin dimethyl ether: CM, drug comparison
  datiscetin: PD, pharmacology
  datiscetin: DT, drug therapy
datiscetin: IT, drug interaction
datiscetin: CM, drug comparison
datiscetin: CB, drug combination
  kaempferol: PD, pharmacology
  kaempferol: DT, drug therapy
kaempferol: IT, drug interaction
kaempferol: CM, drug comparison
kaempferol: CB, drug combination
  oroxylin a: PD, pharmacology
  oroxylin a: DT, drug therapy
oroxylin a: IT, drug interaction
oroxylin a: CM, drug comparison
oroxylin a: CB, drug combination
 primuletin: PD, pharmacology
 primuletin: DT, drug therapy
primuletin: IT, drug interaction
primuletin: CM, drug comparison
primuletin: CB, drug combination
  skullcapflavone ii: PD, pharmacology
  skullcapflavone ii: DT, drug therapy
skullcapflavone ii: IT, drug interaction
skullcapflavone ii: CM, drug comparison
skullcapflavone ii: CB, drug combination
wogonin: IT, drug interaction
  wogonin: DT, drug therapy
```

wogonin: PD, pharmacology
wogonin: CM, drug comparison
wogonin: CB, drug combination

unclassified drug

CAS REGISTRY NO.: (baicalein) 491-67-8; (apigenin) 520-36-5; (bradykinin)

58-82-2, 5979-11-3; (kaempferol) 520-18-3; (oroxylin a) 480-11-5; (skullcapflavone ii) 55084-08-7; (wogonin)

632-85-9

COMPANY NAME: Sigma (United States); Roth (Germany)

L124 ANSWER 32 OF 39 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 89057987 EMBASE

DOCUMENT NUMBER: 1989057987

TITLE: A flavonoid inhibitor of 5-lipoxygenase inhibits

leukotriene production following ischemia in gerbil brain.

AUTHOR: Ban M.; Tonai T.; Kohno T.; Matsumoto K.; Horie T.;

Yamamoto S.; Moskowitz M.A.; Levine L.

CORPORATE SOURCE: Department of Neurological Surgery, School of Medicine,

Tokushima University, Tokushima 770, Japan

SOURCE: Stroke, (1989) 20/2 (248-252). ISSN: 0039-2499 CODEN: SJCCA7

COUNTRY: United States

DOCUMENT TYPE: Journal

FILE SEGMENT: 008 Neurology and Neurosurgery

037 Drug Literature Index

018 Cardiovascular Diseases and Cardiovascular Surgery

026 Immunology, Serology and Transplantation

LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

Leukotrienes C4 and D4 are arachidonic acid metabolites that constrict blood vessels and enhance vascular permeability; their biosynthesis is initiated by the reaction of arachidonic acid with 5-lipoxygenase enzyme. After bilateral carotid artery occlusion for 15 minutes and reperfusion of the gerbil brain for 15 minutes, we determined the brain tissue concentrations of leukotrienes C4 and D4 by radioimmunoassay; they had increased from a baseline concentration of <1 to a mean .+-. SEM concentration of 12.8 .+-. 3.9 pmol/g brain. We also studied the effect of a flavonoid 5-lipoxygenase inhibitor on leukotriene production in the reperfused gerbil brain. A water-soluble flavonoid (5-hexyloxy-3',4'-dihydroxy-6,7-dimethoxyflavone 4'-disodium phoshate) was administered intravenously at a dose of 200 mg/kg body wt; 15 minutes later, both carotid arteries were occluded. The enhanced production of leukotrienes C4 and D4 in the reperfused brain was reduced by approximately 80% (from a mean .+-. SEM of 12.8 .+-. 3.9 to 2.2 .+-. 1.3 pmol/g brain) in the presence of the 5-lipoxygenase inhibitor. The flavonoid did not affect the production of prostaglandin D2, the concentration of which also increased in the reperfused ischemic brain.

CONTROLLED TERM: Medical Descriptors:

*brain ischemia

carotid artery obstruction

gerbil

radioimmunoassay animal experiment

nonhuman

intravenous drug administration

priority journal
Drug Descriptors:

*arachidonate 5 lipoxygenase

*leukotriene

5 hexyloxy 3',4' dihydroxy 6,7 dimethoxyflavone 4'

disodium phosphate: PD, pharmacology

unclassified drug

CAS REGISTRY NO.: (arachidonate 5 lipoxygenase) 80619-02-9

L124 ANSWER 33 OF 39 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER:

2001-476022 [51] WPIDS

DOC. NO. CPI:

C2001-142785

TITLE:

Production of enriched flavonoid aglycone extract for

treating and preventing degenerative diseases,

e.g. heart disease, comprises

enzymatically converting flavonoid glycoside into

flavonoid aglycone, and adjusting acidity.

DERWENT CLASS: B02 D16

INVENTOR(S):
PATENT ASSIGNEE(S):

BURONG, W G; WALLACE, R G (BIOR-N) BIOREX HEALTH LTD

COUNTRY COUNT:

94

PATENT INFORMATION:

PATENT	NO	KIND	DATE	WEEK	LA	PG

WO 2001051482 A1 20010719 (200151)* EN 46

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001026531 A 20010724 (200166)

APPLICATION DETAILS:

PATENT NO K	-113	APPLICATION	DATE
WO 2001051482	 		20010111
AU 2001026531	A	AU 2001-26531	20010111

FILING DETAILS:

PATENT NO	KIND		PATE	ON TH
AU 200102653	1 A	Based on	₩O 2	00151482

PRIORITY APPLN. INFO: US 2000-175443P 20000111; AU 2000-5043

20000111

AB WO 200151482 A UPAB: 20010910

NOVELTY - Producing an enriched flavonoid aglycone extract from starting material containing flavonoid glycoside or its conjugate, comprises enzymatically converting the flavonoid glycoside into flavonoid aglycone, and adjusting the pH to render the flavonoid aglycone soluble, removing the insoluble fraction, and rendering the soluble flavonoid aglycone insoluble.

DETAILED DESCRIPTION - Production of an enriched flavonoid aglycone extract from starting material containing flavonoid glycoside or its conjugate, comprises enzymatically converting the flavonoid glycoside into flavonoid aglycone. The pH is adjusted to render the flavonoid aglycone soluble, removing the insoluble fraction. The pH is adjusted to render the soluble flavonoid aglycone relatively insoluble and forming the flavonoid glycoside extract.

An INDEPENDENT CLAIM is also included for the enriched flavonoid aglycone extract produced by the new method.

ACTIVITY - Antimicrobial; antioxidant; cardiant; neuroprotective; nootropic; cytostatic. No biological data is given.

MECHANISM OF ACTION - None given.

USE - The method is used for the production of enriched flavonoid aglycone (claimed) extract used as therapeutic, anti-microbial, and antioxidant. Flavonoids are used for treating and preventing a range of medical disorders and diseases including degenerative diseases, e.g. heart disease, Alzheimer's disease, dementia, and cancer.

ADVANTAGE - The method does not involve the use of toxic reagents, does not require undue multiple extractions, does not involve extraction of the flavonoid in its glycosylated form (flavonoid glycoside), is not time consuming, and does not involve the use of significant quantities of flammable organic solvents.

Dwq.0/0

L124 ANSWER 34 OF 39 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 2

2001-168573 [17] WPIDS

CROSS REFERENCE:

2000-491047 [41]

DOC. NO. CPI:

C2001-050394

TITLE:

Identifying pattern of cellular responses caused by inhibition of signaling molecule, useful for identifying therapeutic selective inhibitors, particularly of protein

kinases.

DERWENT CLASS:

B04 D16

INVENTOR(S):

BISHOP, A; SHOKAT, K M (UYPR-N) UNIV PRINCETON

PATENT ASSIGNEE(S): COUNTRY COUNT:

95

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001007659 A2 20010201 (200117) * EN 78

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000063620 A 20010213 (200128)

EP 1196626 A2 20020417 (200233) EN

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

APPLICATION DETAILS:

PATENT NO KI	IND	APPLICATION	DATE
WO 2001007659 AU 2000063620 EP 1196626		WO 2000-US19912 AU 2000-63620 EP 2000-950527 WO 2000-US19912	20000721 20000721 20000721 20000721

FILING DETAILS:

PA	TENT NO K	IND			PAT	TENT NO
ΑU	2000063620	Α	Based	on	WO	200107659
ΕP	1196626	A2	Based	on	WO	200107659

PRIORITY APPLN. INFO: US 2000-621293 20000720; US 1999-145422P 19990723

AB WO 200107659 A UPAB: 20020524

NOVELTY - Identifying a pattern of cellular responses attributable to selective inhibition of a particular wild-type signaling molecule (I), is

new.

DETAILED DESCRIPTION - Identifying a pattern of cellular responses attributable to selective inhibition of a particular wild-type signaling molecule (I), is new. Mutant cells (A), having a functionally silent, mutant form of (I), are exposed to a selective inhibitor (II) of the mutant (I), and the cellular responses of (A), before and after exposure, are identified. Optionally, responses are also determined for wild-type cells (A1), unexposed and/or exposed to (II). The observed responses are compared to identify a pattern of responses attributable to selective inhibition of wild-type (I), corresponding to the response pattern attributable to inhibition of mutant (I) in (A).

INDEPENDENT CLAIMS are also included for the following:

- (1) pattern of cellular responses attributable to selective inhibition of wild-type (I), comprising changes in responses to selective inhibition of mutant (I) by (II); and
- (2) identifying a selective inhibitor (IIa) of wild-type (I) by identifying a pattern of responses, treating wild-type cells with test compound and selecting compounds that generate a similar pattern of responses.

ACTIVITY - Cytostatic; vasotropic; antiarteriosclerosis, nephrotropic; antipsoriatic; nootropic; neuroprotective.

No biological data is given.

MECHANISM OF ACTION - (I) inhibitor.

USE - The method is used to establish a pattern of responses that allows identification of selective inhibitors of wild-type (I), particularly protein kinases, from their ability to create a similar response pattern. The selective inhibitors are potentially useful for treating abnormal cell growth, e.g. tumors, restenosis, atherosclerosis, glomerulonephritis, psoriasis and Alzheimer's disease. They can also be used to identify specific substrates and to study biochemical/phenotypic effects of kinase downregulation.

ADVANTAGE - Specific inhibitors of (I) can now be identified without having to express, purify and assay (I). Dwg.0/5

L124 ANSWER 35 OF 39 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER:

2002-138755 [18] WPIDS

CROSS REFERENCE:

1999-263429 [22]; 2001-416769 [38]; 2001-431951 [44]

DOC. NO. CPI:

C2002-042698

TITLE:

Compositions useful in the treatment of

cardiovascular disease e.g.

atherosclerosis and hypercholesterolemia

comprise limonoids e.g. limonin, flavonoids e.g. naringin

and hesperidin and/or tocotrienols e.g. alpha-

tocotrienol.

DERWENT CLASS:

B05

INVENTOR(S):

GUTHRIE, N; KUROWSKA, E M

PATENT ASSIGNEE(S):

(GUTH-I) GUTHRIE N; (KURO-I) KUROWSKA E M

COUNTRY COUNT:

PATENT INFORMATION:

WEEK LA PATENT NO KIND DATE PG -----US 2001055627 A1 20011227 (200218)*

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
US 2001055627 A1 CIP of	US 1997-938640 US 2000-481724	19970926 20000112

FILING DETAILS:

PATENT NO KIND PATENT NO US 2001055627 A1 CIP of US 6251400

PRIORITY APPLN. INFO: US 2000-481724 20000112; US 1997-938640

19970926

US2001055627 A UPAB: 20020319 AΒ

> NOVELTY - A composition (I) comprises a flavonoid selected from hesperidin, naringin, naringenin, hesperitin, nobiletin or tangeretin.

DETAILED DESCRIPTION - A composition (I) comprises a flavonoid selected from hesperidin, naringin, naringenin, hesperitin, nobiletin or tangeretin.

An INDEPENDENT CLAIM is included for a composition (II) comprising a limonoid selected from limonin and nomilin and a tocotrienol.

ACTIVITY - Antiarteriosclerotic; Antilipemic. MECHANISM OF ACTION - Liver cholesterol synthesis inhibitor; low-density lipoprotein (LDL) cholesterol inhibitor.

Rabbits suffering from casein induced hypercholesterolemia were given semi purified cholesterol free casein diet and either water or orange juice. The control group received water to drink and test groups were given orange juice. The different lipoprotein concentration of cholesterol after 3 weeks on test supplement/control was as follows (mg/g liver): total cholesterol = 3.1 plus or minus 0.1/3.8 plus or minus 0.2; cholesterol esters = 0.7 plus or minus 0.1/1.2 plus or minus 0.2; free cholesterol = 2.4 plus or minus 0.1/2.7 plus or minus 0.1. The test supplement reduced the LDL cholesterol levels compared with control. This was associated with significant decrease in liver cholesterol esters but not with increase in fecal excretion of cholesterol and bile acids. The results indicated that the changes in the LDL cholesterol and in liver cholesterol esters might be due to juice components such as limonoids and flavonoids.

USE - The compositions are useful in the treatment of atherosclerosis, hypercholesterolemia (claimed) and hyperlipidemia.

ADVANTAGE - The composition has inhibitory effects on synthesis of liver cholesteryl esters and/or degradation of apo-B proteins. Dwg.0/6

L124 ANSWER 36 OF 39 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 2000-339506 [29] WPIDS

DOC. NO. CPI:

C2000-102976

TITLE:

Increasing plasma beneficial high density lipoprotein levels with bioflavonoids or plant extracts containing them, given as such or in foods and beverages, reduces

risk of coronary disease and atherosclerosis.

DERWENT CLASS:

B02 B03 D13

INVENTOR(S):

AHN, B; BOK, S; CHOI, M; CHOI, Y; HYUN, B; JEONG, T; KIM, S; KWON, Y; LEE, C; LEE, E; LEE, S; MOON, O; MOON, S; AHN, B T; BOK, S H; CHOI, M S; CHOI, Y K; HYUN, B H; JEONG, T S; KIM, S G; KWON, Y K; LEE, C H; LEE, E S; LEE,

S B; MOON, O S; MOON, S S; PARK, Y B

PATENT ASSIGNEE(S):

(KOAD) KOREA ADV INST SCI & TECHNOLOGY

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

23

WO 2000023073 A1 20000427 (200029) * EN 24

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

Page 61

W: CA CN JP RU

US 6133241 A 20001017 (200054)# EP 1123096 A1 20010816 (200147) EN

R: DE FR GB IT

CN 1327384 A 20011219 (200226)

APPLICATION DETAILS:

PATENT NO K	IND	APPLICATION	DATE
WO 2000023073	A1	WO 1998-KR326	19981020
US 6133241	A	US 1998-177448	19981022
EP 1123096	A1	EP 1998-951779	19981020
		WO 1998-KR326	19981020
CN 1327384	A	CN 1998-814276	19981020
		WO 1998-KR326	19981020

FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 1123096	Al Based on	WO 200023073

PRIORITY APPLN. INFO: WO 1998-KR326 19981020; US 1998-177448

19981022

AB WO 200023073 A UPAB: 20000617

NOVELTY - Use of a bioflavanoid (I) or plant extract containing it, for increasing plasma high density lipoprotein levels; and use, as such or in foods and beverages.

DETAILED DESCRIPTION - Use of a bioflavanoid of formula (I), or a plant extract containing it, or of neohesperidin dihydrochalcone of formula (II), for increasing plasma high density lipoprotein levels in a mammal, is new:

---- = an optional bond;

R1-R9 = H, 1-9C alkoxy (optionally substituted by hydroxy, 1-5C alkoxy, aryloxy, or phenyl, 5-9C cycloalkoxy, or 6-10C cycloalkylcarbonyloxy (all optionally substituted by 1-3 Y or amido), 2-10C or 16-18C acyloxy (optionally substituted by hydroxy, 1-5C alkoxy, aryloxy, or phenyl (optionally substituted by Y), or rutinosyl or rhamnosyl; and

Y = hydroxy, alkoxy, aryloxy, halogen, or nitro ACTIVITY - Hypocholesteremic (for HDL cholesterol); cardiovascular. Other activities, reported in prior art, are: antioxidant; anticancer; antiviral; hypotensive.

USE - (I) and (II), and extracts containing them are of value in prevention of cardiovascular disorders linked to low HDL/LDL ratios, notably atherosclerosis.

ADVANTAGE - The bioflavonoids are from natural, rather than synthetic, materials and are non-toxic, even at a level of 1 g/kg. Dwg.0/0

L124 ANSWER 37 OF 39 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 2000-295555 [26] WPIDS

1

DOC. NO. CPI: C2000-089493

TITLE: Medical agent for inhibiting production of matrix

metalloprotease or its precursor, - contains polyalkoxyflavonoid compound, such as nobiletin

or tangeretin.

DERWENT CLASS: B02

PATENT ASSIGNEE(S): (NORQ) NORINSUISANSHO KAJU SHIKENBACHO; (NORQ)

NORINSUISANSHO KAJU SHIKENJOCHO

COUNTRY COUNT:

PATENT INFORMATION:

Page 62

PAT	CENT	NO	KIND	DATE	WEEK	LA	PG
-)210)08003		20000221 20000321	(200026) * (200026)		11 12

APPLICATION DETAILS:

	IND	APPLICATION	DATE
JP 3010210	B1		19980902
JP 2000080035	A	JP 1998-248145	19980902

PRIORITY APPLN. INFO: JP 1998-248145 19980902

AB JP 3010210 B UPAB: 20000606

NOVELTY - A medical agent for inhibiting the production of the matrix metalloprotease or its precursor, contains a polyalkoxyflavonoid. DETAILED DESCRIPTION - A medical agent for inhibiting the production of the matrix metalloprotease or its precursor, contains a polyalkoxyflavonoid of formula (I) R1 = hydrogen or 1-6C alkyl; R2-4 = hydrogen or 1-6C alkoxy; R5 = 1-6C alkyl.

USE - Used in the prevention and/or treatment of matrix metalloprotease-related illnesses, such as chronic rheumatism, osteoarthritis, cancer, arteriosclerosis, aneurysm, cirrhosis, ulcers, osteoporosis, pulmonary fibrosis, glomerulonephritis and peridontal inflammation.

ADVANTAGE - Production of matrix metalloprotease can be inhibited. Dwg.0/8

L124 ANSWER 38 OF 39 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 1999-526388 [44] WPIDS

CROSS REFERENCE: 1999-418245 [35]; 1999-619630 [53]

DOC. NO. CPI: C1999-154680

TITLE: Administering micronutrients and acetylsalicylic acid to

prevent nutritional deficiencies and reduce coronary

heart disease.

DERWENT CLASS: B05

INVENTOR(S): CHRISTAKIS, G; RILEY, P A

PATENT ASSIGNEE(S): (MEDI-N) MEDICAL DOCTORS RES INST INC

COUNTRY COUNT:

PATENT INFORMATION:

PA:	TENT	NO	KIND	DATE	WEEK	LA	PG
US	5948	443	Α	19990907	(199944)*		17

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
US 5948443	A Provisional	US 1996-12158P US 1997-804494	19960223 19970221

PRIORITY APPLN. INFO: US 1996-12158P 19960223; US 1997-804494

AB US 5948443 A UPAB: 19991221

 ${\tt NOVELTY}$ - ${\tt Modular}$ system of multivitamin and mineral supplementation is new.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for a new method to provide micronutrient and acetylsalicylic acid supplementation to treat nutritional deficiencies and to reduce coronary heart

disease in humans comprising the daily administration of a multivitamin/mineral formulation (A) and acetylsalicylic acid.

(A) comprises: Vitamin B1 (0.7-15 mg), vitamin B2 (0.7-15 mg), vitamin B6 (2-100 mg), niacin (6-100 mg), folate (50-800 micro g), pantothenic acid (4-50 mg), vitamin B12 (0.5-40 micro g), biotin (5-300 micro g), calcium (100-1500 mg), magnesium (25-500 mg), iron (1-20 mg), zinc (5-30 mg), manganese (1-10 mg), selenium (10-200 micro g), chromium (10-300 micro g), copper (0-4 mg), Coenzyme Q10 (5-300 mg), vitamin A (200-15000 IU), beta carotene (500-15000 IU), alpha -carotene (50-2000 micro g), lycopene (50-10000 micro g), lutein (50-5000 micro g), zeaxanthin (5-500 micro g), vitamin C (20-1000 mg), vitamin D (0-400 IU), vitamin E (5-2000 mg), grape seed extract (0-300 mg), green tea extract (0-500 mg), crataegus (0-500 mg), oxyacantha extract L-carnitine (0-700 mg), alpha -lipoic acid (0-750 mg), taurine (15-1000 mg), quercitin (0-500 mg) and garlic (0-500 mg).

ACTIVITY - Dietary vitamin supplement; cardiant; antidiabetic; hypotensive; antianemic; cytostatic; osteopathic; antilipemic; thrombolytic; anticoagulant.

A study in seven healthy volunteers compared changes in blood clotting times induced by the modular system (Modules 1 and 4) with the use of conventional multivitamins with acetylsalicylic acid (81 mg). In the two female non-smokers taking the conventional preparations, clotting time was increased from 5.5 to more than 15 minutes. In the three smokers and two non-smokers who took Modules 1 and 4, the clotting times changed from 4-7.5 minutes to 3- more than 15 minutes.

MECHANISM OF ACTION - Platelet deagglutinator; thrombus inhibitor; antioxidant.

Vitamin and antioxidant biochemical action. The combination of acetylsalicylic acid and an antioxidant prevents the oxidation of low density lipoproteins in the coronary artery walls.

USE - For the treatment of nutritional losses and deficiencies and to reduce the risk of coronary **heart disease** (claimed).

To treat or prevent micronutrient deficiency and reduce artherosclerotic-induced coronary heart disease, Syndrome X, diabetes, stress related disorders e.g. mucous colitis and hypertension, immunodeficiency, anemia, fatigue, osteoporosis, cancer, hyperlipidemia and thrombosis in humans. Separate formulations for men and women can be used.

ADVANTAGE - The appropriate formulation matched to specific physiological needs provides optimal results and avoids ingredients counteracting each other or impairing absorption of other ingredients. Platelet deagglutination and thrombus inhibition occur without prolonged blood clotting times and without the side effects of ulceration associated with a higher daily dose of acetylsalicylic acid. Dwg.0/0

L124 ANSWER 39 OF 39 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 1971-67553S [42] WPIDS.

TITLE: 3,3',4',5,7-penta-benzyl-quercetin.

DERWENT CLASS: B02

PATENT ASSIGNEE(S): (LBIO) LABS BIOSEDRA

COUNTRY COUNT: 6

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
BE 765681 JP 46007332	A A		(197142)* (197202)		
DE 2122514 FR 2088127	A A		(197205) (197212)		
GB 1295606 DE 2122514	A B	19740411	(197245)	•	

IT 1036042 B 19791030 (198006)

PRIORITY APPLN. INFO: FR 1970-18458 19700521

AB BE 765681 A UPAB: 19930831

3,3',4',5,7-Penta-benzyl-quercetin Title cpd. useful as a capillary-protecting agent esp. in treating vascular disorders due to arterial hypertension, diabetic and arteriosclerotic, retinitis, chronic glomerulo nephritis hepatic insufficiency, varices of the legs and haemorrhoids are prepd. by benzylating quercitin with benzyl chloride in the presence of KI and K2CO3.

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